=> fil reg; s 77658-84-5
FILE REGTSTRY ENTERED AT 11:14:42 ON 07 JAN 2005
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Property values tagged with IC are from the ZIC/VINITI data file provided by InfoChem.

STRUCTURE FILE UPDATES: 5 JAN 2005 HIGHEST RN 808732-83-4 DICTIONARY FILE UPDATES: 5 JAN 2005 HIGHEST RN 808732-83-4

TSCA INFORMATION NOW CURRENT THROUGH MAY 21, 2004

Please note that search-term pricing does apply when conducting SmartSELECT searches.

Crossover limits have been increased. See HELP CROSSOVER for details.

Experimental and calculated property data are now available. For more information enter HELP PROP at an arrow prompt in the file or refer to the file summary sheet on the web at: http://www.cas.org/ONLINE/DBSS/registryss.html

L4 1 77658-84-5 (77658-84-5/RN)

=> d ide

L4 ANSWER 1 OF 1 REGISTRY COPYRIGHT 2005 ACS on STN

RN 2.7.658=84=5 REGISTRY

CN Benzeneacetamide, N-(2,6-dioxo-3-piperidinyl)- (9CI) (CA INDEX NAME) OTHER NAMES:

CN 3= (Phenylacetylamino) piperidine-2,6-dione-

FS 3D CONCORD

DR 158930-26-8

MF C13 H14 N2 O3

LC STN Files: ADISINSIGHT, ANABSTR, BIOSIS, CA, CAPLUS, IPA, TOXCENTER, USPATFULL

DT.CA CAplus document type: Dissertation; Journal; Patent

RL.P Roles from patents: ANST (Analytical study); BIOL (Biological study); PREP (Preparation); PROC (Process); PRP (Properties); USES (Uses)

RLD.P Roles for non-specific derivatives from patents: BIOL (Biological study)

RL.NP Roles from non-patents: ANST (Analytical study); BIOL (Biological study); PREP (Preparation); PROC (Process); PRP (Properties); USES (Uses)

RLD.NP Roles for non-specific derivatives from non-patents: BIOL (Biological study); USES (Uses)

PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

- 25 REFERENCES IN FILE CA (1907 TO DATE)
- 2 REFERENCES TO NON-SPECIFIC DERIVATIVES IN FILE CA
- 25 REFERENCES IN FILE CAPLUS (1907 TO DATE)

=> => fil capl; d que 121; d que 141; d que 143; d que 142 FILE : CAPLUS ENTERED AT 12:26:46 ON 07 JAN 2005 USE IS SUBJECT TO THE TERMS OF YOUR STN CUSTOMER AGREEMENT. PLEASE SEE "HELP USAGETERMS" FOR DETAILS. COPYRIGHT (C) 2005 AMERICAN CHEMICAL SOCIETY (ACS)

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FILE COVERS 1907 - 7 Jan 2005 VOL 142 ISS 3 FILE LAST UPDATED: 6 Jan 2005 (20050106/ED)

This file contains CAS Registry Numbers for easy and accurate substance identification.

'OBI' IS DEFAULT SEARCH FIELD FOR 'CAPLUS' FILE

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L9
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L10
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L14
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L15
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                                         THREONINE/CN
              2 SEA FILE=REGISTRY ABB=ON VALINE/CN
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L17
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         44310 SEA FILE=CAPLUS ABB=ON (L8 OR L9 OR L10)
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         109418 SEA FILE=CAPLUS ABB=ON (L11 OR L12 OR L13 OR L14 OR L15 OR
L19
               L16)
L20
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L2:1==
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L20 25 SEA FILE=CAPLUS ABB=ON L4
L23 165871 SEA FILE=CAPLUS ABB=ON DRUG DELIVERY SYSTEMS+OLD/CT
L41 2 SEA FILE=CAPLUS ABB=ON L20 AND L23
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L20 25 SEA FILE=CAPLUS ABB=ON L4
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,L43.
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DNA - drug mechanism
of action

1SC.SX - THU- therapeutic use

PKT - pharmacoknetico

BAC - biblogical activity
                <BAC) -/ RL
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             21_SEA_FILE=CAPLUS_ABB=ON_L20_AND_PHARMAC?/SC,SX__,
L42-
=> s 121 or 141 or 143 or 142
∠L127——22—L21—OR—L41—OR—L43=OR=L42—
=> fil uspatf; d que 150; d que 155
FILE USPATFULL ENTERED AT 12:26:47 ON 07 JAN 2005
CA INDEXING COPYRIGHT (C) 2005 AMERICAN CHEMICAL SOCIETY (ACS)
FILE COVERS 1971 TO PATENT PUBLICATION DATE: 6 Jan 2005 (20050106/PD)
FILE LAST UPDATED: 6 Jan 2005 (20050106/ED)
HIGHEST GRANTED PATENT NUMBER: US6839903
HIGHEST APPLICATION PUBLICATION NUMBER: US2005005336
CA INDEXING IS CURRENT THROUGH 6 Jan 2005 (20050106/UPCA)
ISSUE CLASS FIELDS (/INCL) CURRENT THROUGH: 6 Jan 2005 (20050106/PD)
REVISED CLASS FIELDS (/NCL) LAST RELOADED: Oct 2004
USPTO MANUAL OF CLASSIFICATIONS THESAURUS ISSUE DATE: Oct 2004
     USPAT2 is now available. USPATFULL contains full text of the
                                                                            <<<
     original, i.e., the earliest published granted patents or
                                                                            <<<
     applications. USPAT2 contains full text of the latest US
                                                                            <<<
     publications, starting in 2001, for the inventions covered in
                                                                            <<<
     USPATFULL. A USPATFULL record contains not only the original
                                                                            <<<
     published document but also a list of any subsequent
                                                                            <<<
     publications. The publication number, patent kind code, and
                                                                            <<<
     publication date for all the US publications for an invention
                                                                            <<<
     are displayed in the PI (Patent Information) field of USPATFULL
                                                                            <<<
     records and may be searched in standard search fields, e.g., /PN,
                                                                            <<<
     /PK, etc.
                                                                            <<<
     USPATFULL and USPAT2 can be accessed and searched together
                                                                            <<<
     through the new cluster USPATALL. Type FILE USPATALL to
                                                                            <<<
>>>
     enter this cluster.
                                                                            <<<
>>>
                                                                            <<<
>>>
     Use USPATALL when searching terms such as patent assignees,
                                                                            ~~~
>>>
     classifications, or claims, that may potentially change from
                                                                            <<<
>>>
     the earliest to the latest publication.
                                                                            <<<
This file contains CAS Registry Numbers for easy and accurate
substance identification.
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L7
               1 SEA FILE=REGISTRY ABB=ON
                                             RIBOFLAVIN/CN
               2 SEA FILE=REGISTRY ABB=ON
L8
                                             ARGININE/CN
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                                             ORNITHINE/CN
L9
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CITRULLINE/CN

ALANINE/CN

GLYCINE/CN

SERINE/CN

TAURINE/CN

THREONINE/CN

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2 SEA FILE=REGISTRY ABB=ON

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L10

L11

L12 L13

L14

L15

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L50
L4
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            538 SEA FILE=USPATFULL ABB=ON
L52
                T#)/IT
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4703 SEA FILE=USPATFULL ABB=ON (TOXICITY OR CYTOTOXICITY)/IT
6 SEA FILE=USPATFULL-ABB=ON L46-AND (L52-OR-L53)

=> s 150 or 155

L53

L55---

L128 6 L50 OR L55

=> fil biosis; d que 165; d que 168; d que 171

FILE—BIOSIS¹¹ ENTERED AT 12:26:48 ON 07 JAN 2005 Copyright (c) 2005 The Thomson Corporation.

FILE COVERS 1969 TO DATE.

CAS REGISTRY NUMBERS AND CHEMICAL NAMES (CNs) PRESENT FROM JANUARY 1969 TO DATE.

RECORDS LAST ADDED: 5 January 2005 (20050105/ED)

FILE RELOADED: 19 October 2003.

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Ь9
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L11
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                                         GLYCINE/CN
L12
L13
             2 SEA FILE=REGISTRY ABB=ON
                                         SERINE/CN
             1 SEA FILE=REGISTRY ABB=ON
                                          TAURINE/CN
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L15
             2 SEA FILE=REGISTRY ABB=ON VALINE/CN
L16
L58
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L59
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L60
          24253 SEA FILE=BIOSIS ABB=ON
                                       (L8 OR L9 OR L10)
L61
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L62
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L63
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L64
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FILE TOXCENTER ENTERED AT 12:26:51 ON 07 JAN 2005 COPYRIGHT (C) 2005 ACS

FILE COVERS 1907 TO 4 Jan 2005 (20050104/ED)

This file contains CAS Registry Numbers for easy and accurate substance identification.

TOXCENTER has been enhanced with new files segments and search fields. See HELP CONTENT for more information.

TOXCENTER thesauri in the /CN, /CT, and /MN fields incorporate the MeSH 2005 vocabulary. See http://www.nlm.nih.gov/mesh/ and http://www.nlm.nih.gov/pubs/techbull/nd03/nd03_mesh.html for a description of changes. ·

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Ь7
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L8
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                                         ORNITHINE/CN
Ь9
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L10
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                                         ALANINE/CN
L11
             1 SEA FILE=REGISTRY ABB=ON
                                         GLYCINE/CN
L12
            2 SEA FILE=REGISTRY ABB=ON
                                          SERINE/CN
L13
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                                          TAURINE/CN
L14 .
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L15
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L16
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L96
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                                           Ь7
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L99
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                L16)
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T-102
                TAURINE OR THREONINE OR VALINE
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                OR_L101-OR-L102)
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L96
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L104
                T#)
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L105
        2313689 SEA FILE=TOXCENTER ABB=ON TOXIC? OR CYTOTOXIC?
L107
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=> s 1106 or 1108

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=>==>=dup=rem_l127,l128,l92,l129,l130___ FILE 'CAPLUS' ENTERED AT 12:27:28 ON 07 JAN 2005 USE IS SUBJECT TO THE TERMS OF YOUR STN CUSTOMER AGREEMENT.

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L66
         361737 SEA FILE=BIOSIS ABB=ON
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L67
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                                       ((SIDE OR ADVERSE OR TOXIC)(2A)EFFECT#)
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Ļ6.8_
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1.4
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L58
       1857167 SEA FILE=BIOSIS ABB=ON DRUG
L70
       14-SEA-FILE=BIOSIS-ABB=ON-L58-AND-L70-
L71.
=> s 168 or 171
L129 14 L68 OR L71
=> fil ipa; d que 188; d que 192
FILE LIPA' ENTERED AT 12:26:50 ON 07 JAN 2005
COPYRIGHT (C) 2005 American Society of Hospital Pharmacists (ASHP)
FILE COVERS 1970 TO 4 JAN 2005 (20050104/ED)
This file contains CAS Registry Numbers for easy and accurate
substance identification.
              1 SEA FILE=REGISTRY ABB=ON 77658-84-5
1.4
              1 SEA FILE=REGISTRY ABB=ON RIBOFLAVIN/CN
L7
              2 SEA FILE=REGISTRY ABB=ON ARGININE/CN
L<sub>8</sub>
              2 SEA FILE=REGISTRY ABB=ON ORNITHINE/CN
Ь9
             1 SEA FILE=REGISTRY ABB=ON CITRULLINE/CN
L10
             2 SEA FILE=REGISTRY ABB=ON ALANINE/CN
L11
             1 SEA FILE=REGISTRY ABB=ON GLYCINE/CN
L12
             2 SEA FILE=REGISTRY ABB=ON SERINE/CN
L13
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L14
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L15
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L16
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L81
           223 SEA FILE=IPA ABB=ON L7
L82
           291 SEA FILE=IPA ABB=ON RIBOFLAVIN OR VITAMIN(W)(B2 OR B 2)
L83
           149 SEA FILE=IPA ABB=ON (L8 OR L9 OR L10)
L84
L85
           588 SEA FILE=IPA ABB=ON ARGININE OR ORNITHINE OR CITRULLINE
           212 SEA FILE=IPA ABB=ON (L11 OR L12 OR L13 OR L14 OR L15 OR L16)
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1.87
                OR THREONINE OR VALINE
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                OR-L87-)----
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L81
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L90
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L91
        6-SEA-FILE=IPA-ABB=ON-L81-AND-(L89-OR-L90-OR-L91-)
L92=
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=> fil toxcenter; d que 1106; d que 1108

Page 7

Jones PLEASE SEE "HELP USAGETERMS" FOR DETAILS. COPYRIGHT (C) 2005 AMERICAN CHEMICAL SOCIETY (ACS) FILE 'USPATFULL' ENTERED AT 12:27:28 ON 07 JAN 2005 CA INDEXING COPYRIGHT (C) 2005 AMERICAN CHEMICAL SOCIETY (ACS) FILE 'IPA' ENTERED AT 12:27:28 ON 07 JAN 2005 COPYRIGHT (C) 2005 American Society of Hospital Pharmacists (ASHP) FILE 'BIOSIS' ENTERED AT 12:27:28 ON 07 JAN 2005 Copyright (c) 2005 The Thomson Corporation. FILE 'TOXCENTER' ENTERED AT 12:27:28 ON 07 JAN 2005 COPYRIGHT (C) 2005 ACS PROCESSING COMPLETED FOR L127 PROCESSING COMPLETED FOR L128 PROCESSING COMPLETED FOR L92 PROCESSING COMPLETED FOR L129 PROCESSING COMPLETED FOR L130 L131 44-DUP_REM_L127_L128-L92-L129_L130 (14-DUPLICATES_REMOVED) 1 ANSWERS '1-22' FROM FILE CAPLUS ANSWERS '23-27' FROM FILE USPATFULL ANSWERS '28-32' FROM FILE IPA ANSWERS '33-43' FROM FILE BIOSIS ANSWER '44' FROM FILE TOXCENTER =>dibibedabhitind1=22;dibibabhitrn23-27;-dial1-28-44 L131 ANSWER 1 OF 44 CAPLUS COPYRIGHT 2005 ACS on STN DUPLICATE 1 2003:434352 CAPLUS ACCESSION NUMBER: 138:406977 DOCUMENT NUMBER: Formulation of amino acids and riboflavin useful to TITLE: reduce toxic effects of cytotoxic chemotherapy INVENTOR(S): Burzynski, Stanislaw R. PATENT ASSIGNEE(S): USA PCT Int. Appl., 25 pp. SOURCE: CODEN: PIXXD2 DOCUMENT TYPE: Patent LANGUAGE: English FAMILY ACC. NUM. COUNT: PATENT INFORMATION: APPLICATION NO. PATENT NO. KIND DATE DATE ----------_ _ _ _ _ _ _ _ WO 2002-US3 WO 2003045372 A1 20030605 **V**354 20021121 W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ÆS, A, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW/MX, MZ, NO, NZ, OM, PH, PL, PT, RO, RU, SC, SD, SE, SG, SI, SK, SL, TJ, My, TN, TR, TT, TZ, UA, UG, UZ, VC, VN, YU, ZA, ZM, ZW

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RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, 7/Z, UG, ZM, ZW, AM, AZ, BY,
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             FI, FR, GB, GR, IE, IT, LU, MC, NL,/PT, SE, SK, TR, BF, BJ, CF,
             CG, CI, CM, GA, GN, GQ, GW, ML, MR/ NE, SN, TD, TG
                                            US 2001-995010
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            AT, BE, CH, DE, DK, ES, FR, GB, ÇR, IT, LI, LU, NL, SE, MC, PT,
             IE, SI, LT, LV, FI, RO, MK, CY, AL, TR, BG, CZ, EE, SK
                                            BR 2002-14430
     BR 2002014430
                         Α
                                20041103
                                                                    20021121
PRIORITY APPLN. INFO.:
                                            US 2001-995010
                                                                A 20011127
                                            WO 2002-US37354
                                                               W 20021121
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ED Entered STN: 06 Jun 2003

AB Pharmaceutical compns. effective in alleviating or reducing the effects of fatigue and weakness associated with cancer and cytotoxic cancer chemotherapy are disclosed. The pharmaceutical compns. of the present invention comprise riboflavin, effectors of the urea cycle in free form or pharmacol. acceptable salts thereof, and amino acids selected from the groups of essential and non-essential amino acids, in free form or pharmaceutically acceptable salts thereof, suitably combined with appropriate carriers, diluents, or excipients. Also disclosed are methods of alleviating or reducing the effects of fatigue and weakness associated with cancer and cytotoxic cancer chemotherapy by administration of pharmaceutical compns. of the present invention.

IC ICM A61K031-195

ICS A61K031-198; A61K031-525; A61P041-00; A23L001-305

CC 63-6 (Pharmaceuticals)

Section cross-reference(s): 1

IT Drug delivery systems

(carriers; formulation of amino acids and riboflavin useful to reduce toxic effects of cytotoxic chemotherapy)

IT Drug delivery systems

(injections, i.v.; formulation of amino acids and riboflavin useful to reduce toxic effects of cytotoxic chemotherapy)

IT Drug delivery systems

(parenterals; formulation of amino acids and riboflavin useful to reduce toxic effects of cytotoxic chemotherapy)

IT 56-40-6, Glycine, biological studies 56-41-7, Alanine,
biological studies 56-45-1, Serine, biological studies
70-26-8, Ornithine 72-18-4, Valine, biological studies
72-19-5, Threonine, biological studies 74-79-3,

Arginine, biological studies 83-88-5, Riboflavin, biological

studies 107-35-7, Taurine 372-75-8, Citrulline

77658-84-5, 3-Phenylacetylamino-2,6-piperidinedione

RL: PAC (Pharmacological activity); PEP (Physical, engineering or chemical process); PYP (Physical process); THU (Therapeutic use); BIOL (Biological study); PROC (Process); USES (Uses)

(formulation of amino acids and riboflavin useful to reduce toxic effects of cytotoxic chemotherapy)

REFERENCE COUNT:

THERE ARE 4 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L131 ANSWER 2 OF 44 CAPLUS COPYRIGHT 2005 ACS on STN DUPLICATE 4

ACCESSION NUMBER:

1998:31104 CAPLUS

DOCUMENT NUMBER:

128:111158

TITLE:

Design of drugs involving receptor-ligand-DNA

interactions

INVENTOR(S):

Hendry, Lawrence B.

PATENT ASSIGNEE(S):

USA

SOURCE:

U.S., 22 pp., Cont.-in-part of U.S. Ser. No. 158,689.

CODEN: USXXAM

DOCUMENT TYPE:

Patent

LANGUAGE:

English

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
US 5705335	A	19980106	US 1994-369779	19941128
US 5888738	Α	19990330	US 1997-864669	19970528
US 5888741	Α	19990330	US 1997-935219	19970822
US 6306595	B1	20011023	US 1999-239491	19990128
US 2002064790	A1	20020530	US 2001-941230	20010828
PRIORITY APPLN. INFO.:			US 1993-158689	A2 19931126

US 1994-369779 A1 19941128 US 1997-864669 A1 19970528 US 1999-239491 A1 19990128

ED Entered STN: 19 Jan 1998

It has been discovered that the degree of hormonal activity of candidate AR ligands correlates better with the degree of fit into DNA than with the strength of receptor binding, and that the receptors in the steroid/thyroid hormone/vitamin A and D family alter the physiochem. properties of DNA and in concert with other transcription factors facilitate insertion of the ligand into DNA. As a result, the magnitude of the response is a function of the structure of the ligand as it relates to insertion and fit into the DNA and the specificity of the response is a function of the stereochem. of the receptor through binding to both the ligand and to the DNA. Based on these discoveries, a computer-based method is described herein for identifying drugs having increased activity as compared with the natural ligand for receptors such as the estrogenic receptors.

IC ICM C12Q001-68

ICS G06F017-50; A61K031-56

NCL 435006000

CC 2-2 (Mammalian Hormones)

TT 50-28-2, Estradiol, biological studies 57-83-0, Progesterone, biological 77-06-5, Gibberellic acid 128-20-1 516-54-1, 3α -Hydroxy- 5α -pregnan-20-one 521-18-6, 5α -

Dihydrotestosterone 571-22-2, 5β-Dihydrotestosterone 5864-38-0 77658-84-5, 3-Phenylacetylamino-2,6-piperidinedione 95258-51-8 RL: BAC (Biological activity or effector, except adverse); BPR (Biological process); BSU (Biological study, unclassified); PRP

(Properties); BIOL (Biological study); PROC (Process)

(design of drugs involving receptor-ligand-DNA interactions)

THERE ARE 38 CITED REFERENCES AVAILABLE FOR THIS REFERENCE COUNT: 38 RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L131 ANSWER 3 OF 44 CAPLUS COPYRIGHT 2005 ACS on STN DUPLICATE 5

ACCESSION NUMBER: 1994:499064 CAPLUS

DOCUMENT NUMBER: 121:99064

TITLE: Antiestrogenic piperidinediones designed prospectively

using computer graphics and energy calculations of

DNA-ligand complexes

Hendry, Lawrence B.; Chu, Chung K.; Copland, John A.; AUTHOR (S):

Mahesh, Virendra B.

CORPORATE SOURCE: Dep. Physiol. Endocrinol., Med. Coll. Georgia,

Augusta, GA, 30912, USA

SOURCE: Journal of Steroid Biochemistry and Molecular Biology

(1994), 48(5-6), 495-505 CODEN: JSBBEZ; ISSN: 0960-0760

DOCUMENT TYPE: Journal LANGUAGE: English ED Entered STN: 03 Sep 1994

Drug design technol. based upon DNA stereochem. and now supplemented by AB computer modeling was used to design a novel compound to inhibit estrogen-induced tumor cell growth. A known compound 3-phenylacetylamino-2,6-piperidinedione (PP) was accommodated in partially unwound DNA in a manner consistent with criteria for antiestrogens. Examination of the PP-DNA complex revealed that substitution of a hydroxyl group at the para position (p-OH-PP) would provide a stereospecific hydrogen bond and a substantial increase in fit as assessed by energy calcns. antiestrogen tamoxifen could also be accommodated within the site; analogous substitution of a hydroxyl at the 4-position resulted in a better fitting mol. 4-Hydroxytamoxifen is a more potent antiestrogen than tamoxifen. Synthesis and subsequent evaluation of p-OH-PP as an inhibitor of estrogen stimulated MCF-7 (E3) human breast cancer cell growth

demonstrated that p-OH-PP was more active than both PP and its hydrolysis product phenylacetylglutamine. As predicted, the order of fit into DNA correlated with the relative ability to inhibit estrogen-induced growth of tumor cells suggesting that the evolving drug design technol. will be valuable in developing new drugs for breast cancer.

CC 1-3 (Pharmacology)

Section cross-reference(s): 2

IT 50-28-2, Estradiol, biological studies 10540-29-1, Tamoxifen 28047-15-6, Phenylacetylglutamine 68047-06-3, 4-Hydroxytamoxifen 77658-84-5, 3-Phenylacetylamino-2,6-piperidinedione

RL: BIOL (Biological study)

(DNA binding by, computer modeling of, in antiestrogenic neoplasm inhibitor development)

L131 ANSWER 4 OF 44 CAPLUS COPYRIGHT 2005 ACS on STN DUPLICATE 6

ACCESSION NUMBER: 1994:23166 CAPLUS

DOCUMENT NUMBER: 120:23166

TITLE: Inhibition of estrogen stimulated mitogenesis by

3-phenylacetylamino-2,6-piperidinedione and its

para-hydroxy analog

AUTHOR(S): Copland, John A.; Hendry, Lawrence B.; Chu, Chung K.;

Wood, Joseph C.; Wrenn, Robert W.; Pantazis, Cooley

G.; Mahesh, Virendra B.

CORPORATE SOURCE: Dep. Physiol. Endocrinol., Med. Coll. Georgia,

Augusta, GA, 30912-3000, USA

SOURCE: Journal of Steroid Biochemistry and Molecular Biology

(1993), 46(4), 451-62

CODEN: JSBBEZ; ISSN: 0960-0760

DOCUMENT TYPE: Journal LANGUAGE: English ED Entered STN: 22 Jan 1994

AB 3-Phenylacetylamino-2,6-piperidinedione (A10) inhibited estradiol-stimulated cell growth in the MCF-7 (E3) human breast tumor cell line in vivo and in vitro. While high concns. of A10 were needed to inhibit cell proliferation (IC50 = 3 + 10-3M in vitro), the compound demonstrated little toxicity. The effect appeared specific since a hydrolysis product of A10, phenylacetylglutamine, demonstrated no growth inhibitory activity at similar concns. in MCF-7 (E3) cells in vitro. A computer designed analog, p-hydroxy A10, was more potent than A10 in inhibiting activity in MCF-7 (E3) cells in vitro. The IC50 for p-hydroxy AlO was 7 + 10-6 M which was comparable to that of the antiestrogen, tamoxifen (IC50 1 + 10-7 M). All three compds. caused a decline in estrogen receptor levels in a dose-dependent fashion. Al0 also inhibited estradiol induction of progesterone receptors. Examination of protein kinase activity following an acute exposure to a 10-11 M growth stimulatory dose of estradiol revealed a 168% increase in protein kinase activity over that of untreated control cells. AlO in a dose-responsive fashion inhibited the estradiol-stimulated increase in protein kinase activity. The protein kinase activity was also inhibited by p-hydroxy AlO. These activities of AlO and p-hydroxy AlO coupled with the low toxicity and novelty of the basic A10 structure provide an exciting possibility of developing a new class of clin. useful antineoplastic drugs with minimal side effects.

CC 1-6 (Pharmacology)

Section cross-reference(s): 2

IT 77658-84-5, 3-Phenylacetylamino-2,6-piperidinedione 138592-72-0

RL: BIOL (Biological study)

(breast cancer cells of human inhibition by)

L131 ANSWER 5 OF 44 CAPLUS COPYRIGHT 2005 ACS on STN DUPLICATE 10

ACCESSION NUMBER: 1984:416929 CAPLUS

DOCUMENT NUMBER: 101:16929

TITLE: Animal toxicology studies on oral formulation of

antineoplaston A10

AUTHOR(S): Burzynski, S. R.; Mohabbat, M. O.; Burzynski, B. CORPORATE SOURCE: Burzynski Res. Inst., Stafford, TX, 77477, USA

SOURCE: Drugs under Experimental and Clinical Research (1984),

10(2), 113-18

CODEN: DECRDP; ISSN: 0378-6501

DOCUMENT TYPE: Journal LANGUAGE: English ED Entered STN: 21 Jul 1984

Antineoplaston A10 (I) [77658-84-5] tested in tissue culture of human carcinoma of the breast line MDA-MB-231, produced a cytostatic effect at 2 mg/mL. LD50 in mice of the Na salt of Antineoplaston A10 given in the form of i.p. injection was determined as 10.33 g/kg. In chronic toxicity studies, Antineoplaston A10 was given to the mice in the form of 1.0%, 1.5%, and 2.0% food mixture daily for 30 days and in the form of 1.0% food mixture daily for 365 days. The dosage levels were approx. 1.0 g/kg/day of Antineoplaston A10 for 1.0% food mixture, 1.25 g/kg/day for 1.5%, and 1.5 g/kg/day for 2.0%. A total of 160 mice were used in the expts. The animals were sacrificed on days 30, 60, 90, 180, and 365 and underwent complete phys., gross pathol., and microscopic examination The studies did not reveal any toxic effect associated with daily chronic oral administration of Antineoplaston A10 to mice.

CC 1-6 (Pharmacology)

IT 77658-84-5

RL: PRP (Properties)

(toxicity of, as neoplasm inhibitor, in human cells and laboratory animals)

L131 ANSWER 6 OF 44 CAPLUS COPYRIGHT 2005 ACS on STN

ACCESSION NUMBER: 2004:1036404 CAPLUS

DOCUMENT NUMBER: 142:11305

TITLE: Toothpaste containing anticancer agents and calcium

salts

INVENTOR(S): Burzynski, Stanislaw R.; Gruszecki, Wojciech

PATENT ASSIGNEE(S): USA

SOURCE: U.S. Pat. Appl. Publ., 8 pp.

CODEN: USXXCO

DOCUMENT TYPE: Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
US 2004241107	A1	20041202	US 2003-446536	20030528
PRIORITY APPLN. INFO.:			US 2003-446536	20030528

ED Entered STN: 03 Dec 2004

AB A novel dentifrice composition is provided for prevention or treatment of carcinoma of the oral cavity, caries and periodontal diseases of the oral cavity. The dentifrice composition contains a partially water-soluble calcium salt, a medicinal composition useful in the treatment of human neoplastic disease, and a hydrophilic or hydrophobic liquid vehicle. A preferred dentifrice composition is a toothpaste comprising gypsum, 3-N-phenylacetylamino-2,6-piperidinedione, gypsum, paraffin oil and a mixture of natural flavoring oils. The components of the dentifrice composition act advantageously to allow the composition to remove plaque, tartar, and oral disease-causing bacteria.

IC ICM A61K007-16

NCL 424049000

CC 62-7 (Essential Oils and Cosmetics)
 Section cross-reference(s): 63

IT 77658-84-5

RL: COS (Cosmetic use); PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(toothpaste containing anticancer agents and calcium salts) L131 ANSWER 7 OF 44 CAPLUS COPYRIGHT 2005 ACS on STN 2000:84607 CAPLUS ACCESSION NUMBER: DOCUMENT NUMBER: 132:132326 Antitumor regimen for administration of TITLE: phenylacetylglutamine, phenylacetylisoglutamine, and/or phenylacetate INVENTOR(S): Burzynski, Stanislaw R. PATENT ASSIGNEE(S): USA PCT Int. Appl., 42 pp. SOURCE: CODEN: PIXXD2 DOCUMENT TYPE: Patent LANGUAGE: English FAMILY ACC. NUM. COUNT: PATENT INFORMATION: KIND DATE PATENT NO. APPLICATION NO. DATE _____ _____ _ _ _ _ _____ _____ WO 1999-US15017 19990702 WO 2000004894 A2 20000203 WO 2000004894 А3 20000427 W: AE, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CU, CZ, DE, DK, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN, MW, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, UA, UG, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM RW: GH, GM, KE, LS, MW, SD, SL, SZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG US 1998-121567 US 6258849 В1 20010710 19980723 MX 1999-255 CA 1999-2336945 MX 9900255 Α 20000731 19990104 AA A1 CA 1999-2336945 CA 2336945 20000203 19990702 AU 1999-48542 AU 9948542 20000214 19990702 B2 AU 759278 20030410 Α BR 1999-12356 BR 9912356 20010417 19990702 A2 EP 1999-932179 EP 1098643 20010516 19990702 В1 EP 1098643 20040107 R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO

 JP 2002521329
 T2
 20020716
 JP 2000-560887
 19990702

 AT 257378
 E
 20040115
 AT 1999-932179
 19990702

 NZ 509244
 A
 20040227
 NZ 1999-509244
 19990702

NZ 509244 A 20040227 NZ 1999-509244 19990702 ES 2214866 T3 20040916 ES 1999-932179 19990702

US 2001044466 A1 20011122 US 2001-863035 20010522 PRIORITY APPLN. INFO.: US 1998-121567 A 19980723

OTHER SOURCE(S): MARPAT 132:132326

ED Entered STN: 04 Feb 2000

AB A method of treating neoplastic disease, including cancer, comprises administering a pharmaceutical composition comprising a highly concentrated aqueous solution

of phenylacetylglutamine and phenylacetylisoglutamine in a 4:1 ratio, at an infusion rate of from 100 mL/h to 400 mL/h. Also, the method comprises a highly concentrated aqueous solution of phenylacetate and (phenylacetylglutamine or phenylacetylisoglutamine) in a 4:1 ratio, at an infusion rate of from 100 mL/h to 400 mL/h.

WO 1999-US15017

W 19990702

IC ICM A61K031-00

CC 1-6 (Pharmacology)

Section cross-reference(s): 63

IT 114-70-5, Sodium phenylacetate 1821-12-1, Benzenebutanoic acid 28047-15-6, Phenylacetylglutamine 77658-84-5,

09/955010 Page 13 Jones

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3-Phenylacetylamino-2,6-piperidinedione
                                               104771-88-2
     RL: BAC (Biological activity or effector, except adverse); BSU
     (Biological study, unclassified); THU (Therapeutic use); BIOL
     (Biological study); USES (Uses)
        (antitumor regimen for administration of phenylacetylglutamine,
        phenylacetylisoglutamine, and/or phenylacetate)
L131 ANSWER 8 OF 44 CAPLUS COPYRIGHT 2005 ACS on STN
ACCESSION NUMBER:
                         1999:714113 CAPLUS
DOCUMENT NUMBER:
                         132:54936
TITLE:
                         Enantioselective separation of several
                         piperidine-2,6-dione drugs on Chirose C-1 chiral
                         stationary phase
                         Aboul-Enein, Hassan Y.; Al-Duraibi, Ibrahim A.
AUTHOR(S):
CORPORATE SOURCE:
                         Bioanalytical and Drug Development Laboratory
                         Biological and Medical Research Department (MBC 03),
                         King Faisal Specialist Hospital and Research Centre,
                         Riyadh, 11211, Saudi Arabia
                         Separation Science and Technology (1999), 34(15),
SOURCE:
                         2973-2979
                         CODEN: SSTEDS; ISSN: 0149-6395
                         Marcel Dekker, Inc.
PUBLISHER:
                         Journal
DOCUMENT TYPE:
                         English
LANGUAGE:
     Entered STN: 09 Nov 1999
     A newly developed Chirose C-1 chiral stationary phase, a highly chiral
     polymer, was used for direct and isocratic enantiomeric separation of 12
     piperidine-2,6-dione compds. under normal phase conditions. Baseline
     separation was achieved for 8 compds., 3 compds. were partially separated, while 1
     compound did not resolve.
     64-3 (Pharmaceutical Analysis)
     50-35-1, Thalidomide
                            841-67-8
                                       2614-06-4
                                                   17575-58-5
                                                                17575-59-6
                               57288-03-6 77658-84-5 83155-00-4
     38473-28-8 55511-44-9
                                                            119095-49-7
                              108816-41-7 108929-54-0
     91531-30-5
                  108816-40-6
                                 123979-23-7
                 121742-47-0
                                               123979-24-8
                                                             124095-07-4
     121742-46-9
                  124095-09-6
                                 124095-10-9
                                               137623-89-3
                                                             137623-91-7
     124095-08-5
                                 252943-15-0
                                               252943-17-2
                                                             252943-18-3
                  162662-87-5
     151410-41-2
     RL: ANT (Analyte); ANST (Analytical study)
        (enantioselective separation of piperidinedione drugs on Chirose C-1 chiral
        stationary phase)
                               THERE ARE 26 CITED REFERENCES AVAILABLE FOR THIS
REFERENCE COUNT:
                         26
                               RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT
L131 ANSWER 9 OF 44 CAPLUS COPYRIGHT 2005 ACS on STN
                         1998:350328 CAPLUS
ACCESSION NUMBER:
                         129:48960
DOCUMENT NUMBER:
                         Enantiomeric separation of several cyclic imides on a
TITLE:
                         macrocyclic antibiotic (vancomycin) chiral stationary
                         phase under normal and reversed phase conditions
AUTHOR (S):
                         Aboul-Enein, Hassan Y.; Serignese, Vince
                         Bioanalytical and Drug Development Laboratory,
CORPORATE SOURCE:
                         Biological and Medical Research (MBC-03), King Faisal
                         Specialist Hospital and Research Centre, Riyadh,
                         11211, Saudi Arabia
                         Chirality (1998), 10(4), 358-361
SOURCE:
                         CODEN: CHRLEP; ISSN: 0899-0042
PUBLISHER:
                         Wiley-Liss, Inc.
DOCUMENT TYPE:
                         Journal
                         English
LANGUAGE:
     Entered STN: 10 Jun 1998
     Several cyclic imidic compds. (barbiturates, piperidine-2,6-diones, and
```

ED

AB

CC IT

ED

AB

mephenytoin) are enantiomerically resolved via HPLC on a macrocyclic

antibiotic covalently bonded to a silica gel support. The Chirobiotic V chiral stationary phase (CSP) column contains the antibiotic vancomycin as the chiral selector. The results of the anal. show that the substituents at the chiral carbon position of the racemic drugs affect chiral resolution Ring size may also play a role when considering the formation of analyte-CSP inclusion complexes. Contrary to the piperidine-2,6-diones, the chromatog. parameters for the barbiturates are much the same under normal- or reversed-phase conditions. The details of these results are discussed.

CC 80-4 (Organic Analytical Chemistry)
Section cross-reference(s): 64

50-12-4, (\pm) -Mephenytoin 50-35-1, (\pm) -Thalidomide 56 (\pm) -Hexobarbital 77-21-4, (\pm) -Glutethimide 115-38-8, (\pm) -Mephobarbital 125-84-8, (\pm) -Aminoglutethimide 84 56-29-1, IT 841-67-8, (-)-Thalidomide 1156-05-4, (±)-Phenglutarimide 2303-80-2, 2671-99-0, 2614-06-4, (+)-Thalidomide (-)-Mephobarbital 4336-84-9, (\pm) -1,5-Dimethyl-5-phenylbarbituric acid (+)-Mephobarbital 7245-06-9, (-)-Hexobarbital 17575-58-5, 7245-04-7, (+)-Hexobarbital 17575-59-6, (-)-Glutethimide (+)-Glutethimide 28900-81-4, (-)-1,5-Dimethyl-5-phenylbarbituric acid 28900-82-5, (+)-1,5-Dimethyl-5-phenylbarbituric acid 36045-93-9, (±)-1-Methyl-5-phenyl-5-n-propylbarbituric acid 37120-83-5, (-)-1-Methyl-5-phenyl-5-n-propylbarbituric acid 37120-84-6, (+)-1-Methyl-5-phenyl-5-n-propylbarbituric acid 55511-44-9, (+)-Aminoglutethimide 57288-03-6, (-)-Aminoglutethimide 70989-04-7. 71140-51-7, (-)-Mephenytoin 77658-84-5 (+)-Mephenytoin 91531-30-5 92788-10-8, (±)-Pyridoglutethimide 108929-54-0 112798-45-5, (+)-Phenglutarimide 112924-18-2, (-)-Phenglutarimide 121742-46-9, (+)-Pyridoglutethimide 121742-47-0, (-)-Pyridoglutethimide RL: ANT (Analyte); PEP (Physical, engineering or chemical process); ANST

(Analytical study); PROC (Process)

(enantiomeric separation of several cyclic imides by HPLC on Chirobiotic V chiral stationary phase under normal and reversed phase conditions)

REFERENCE COUNT: 10 THERE ARE 10 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L131 ANSWER 10 OF 44 CAPLUS COPYRIGHT 2005 ACS on STN

ACCESSION NUMBER:

1997:752824 CAPLUS

DOCUMENT NUMBER:

128:39559

TITLE:

Liposomal antineoplaston therapies with markedly

improved antineoplastic activity

INVENTOR(S):

Byra, Anna R.; Burzynski, Stanislaw R.; Waldbillig,

Robert J.

PATENT ASSIGNEE(S):

Burzynski Research Institute, USA

SOURCE:

PCT Int. Appl., 40 pp. CODEN: PIXXD2

DOCUMENT TYPE:

Patent

LANGUAGE:

English

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

PATENT NO.	KIND DATE	APPLICATION NO.	DATE
WO 9742939	A1 19971120	WO 1997-US8167	19970514
W: CA, JP, KR			
RW: AT, BE, CH,	DE, DK, ES, FI,	FR, GB, GR, IE, IT, LU,	MC, NL, PT, SE
CA 2254772	AA 19971120	CA 1997-2254772	19970514
CA 2254772	C 20040127		
EP 906088	A1 19990407	EP 1997-923650	19970514
R: AT, BE, CH,	DE, DK, ES, FR,	GB, GR, IT, LI, LU, NL,	SE, MC, PT,
IE, FI			
US 6013278	A 20000111	US 1997-856133	19970514

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JP 2002503209
                        T2
                              20020129
                                          JP 1997-541103
                                                                19970514
                                                           P 19960514
PRIORITY APPLN. INFO.:
                                          US 1996-17616P
                                          US 1997-856133
                                                            A 19970514
                                          WO 1997-US8167
                                                            W 19970514
```

ED Entered STN: 03 Dec 1997

AB A second generation of antineoplaston therapies with markedly improved antineoplastic activity is disclosed. Among others, members of the antineoplaston family include phenylacetate (PN), 3-phenylacetylamino-2,6piperidinedione (CN), and hydrolysis derivs. of CN: phenylacetylglutamine (PG) and isophenylacetylglutamine (Iso-PG). In part, these increases in antineoplastic activity result from large increases in the transport of antineoplaston compns. into cells. Importantly and unexpectedly these increases in antineoplastic activity also result from the capacity of the drug delivery system to direct antineoplaston compds. intracellular trafficking to intracellular binding sites influencing cell viability and proliferation. Liposomal formulations of antineoplaston compns. increase in vitro antineoplastic activity by a factor of 750 to 1500 as compared to administration of antineoplaston compds. given without liposomal formulations. In addition, these liposomal formulations enhanced cellular uptake of antineoplaston compds. form 30 to more that 80 fold. Liposomal formulations were also found to increase intracellular levels of the antineoplaston CN by directing CN to intracellular binding sites that influence cell viability and proliferation and block its hydrolysis. Under conditions where free CN has no antineoplastic activity, liposomally formulated CN can produce essentially complete and relatively long-lasting blockade of cell growth. Cell growth was found to be restored as intracellular levels of bound CN decrease to undetectable levels.

TC ICM A61K009-127

ICS A61K031-445; A61K031-195; A61K031-19

63-6 (Pharmaceuticals) CC

Section cross-reference(s): 1

IT Drug delivery systems

(liposomes; liposomal antineoplaston therapies)

103-82-2, Phenylacetic acid, biological studies 28047-15-6, TT Phenylacetylglutamine 77658-84-5, 3-Phenylacetylamino-2,6piperidinedione 184849-22-7

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); PEP (Physical, engineering or chemical process); THU (Therapeutic use); BIOL (Biological study); PROC (Process); USES (Uses)

(liposomal antineoplaston therapies)

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L131 ANSWER 11 OF 44 CAPLUS COPYRIGHT 2005 ACS on STN
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ACCESSION NUMBER: 1997:639120 CAPLUS

DOCUMENT NUMBER:

127:341205

TITLE:

Enantioselective separation of several

piperidine-2,6-diones on a covalently bonded cellulose

3,5-dimethylphenyl carbamate/10-undecenoate chiral

selector

AUTHOR(S):

Aboul-Enein, Hassan Y.; Serignese, Vince; Minguillon,

Cristina; Oliveros, Laureano

CORPORATE SOURCE:

Bioanalytical Drug Development Lab., Biol. Medical Res. (MBC-03), King Faisal Specialist Hospital Res.

Centre, Riyadh, 11211, Saudi Arabia

SOURCE:

Biomedical Chromatography (1997), 11(5), 303-306

CODEN: BICHE2; ISSN: 0269-3879

PUBLISHER: Wiley DOCUMENT TYPE: Journal LANGUAGE: English

ED Entered STN: 09 Oct 1997

A series of piperidine-2,6-dione-based drugs were enantiomerically AB resolved on a covalently bonded cellulose 3,5-dimethylphenyl

09/955010 Jones Page 16

carbamate/10-undecenoate chiral stationary phase (CSP), under normal- or reversed-phase conditions. A single column that can be applied in both modes may be advantageous when considering the shorter setup time required and the solubility of the compound to be analyzed since many samples possess different solubilities. The covalently bonded CSP was compared to a com. available chiral adsorbent, Chiralcel OD, which was previously used for the enantiomeric resolution of the above-mentioned drug series. Chiralcel OD was used in the normal-phase mode and was more fragile than the column used here. In addition, the range of solvents available as eluents was more restricted. Thus, it was of interest to look at the possible advantages of using a chemical bonded phase that can be applied in dual mode.

CC 80-4 (Organic Analytical Chemistry) Section cross-reference(s): 27, 63

77-21-4, Glutethimide 125-84-8, Aminoglutethimide TT 1121-89-7D, 2,6-Piperidinedione, chiral derivs. 38473-28-8, Acetylaminoglutethimide 77658-84-5 115883-22-2

RL: ANT (Analyte); ANST (Analytical study)

(enantiomeric separation of; enantioselective separation of piperidine-2,6-dione pharmaceuticals with covalently bonded cellulose stationary phase)

THERE ARE 10 CITED REFERENCES AVAILABLE FOR THIS REFERENCE COUNT: 10

RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L131 ANSWER 12 OF 44 CAPLUS COPYRIGHT 2005 ACS on STN

1997:222135 CAPLUS ACCESSION NUMBER:

DOCUMENT NUMBER: 126:347362

Enantiomeric separation of some piperidine-2,6-dione TITLE:

drugs using chiralcel OJ-R column

Aboul-Enein, Hassan Y.; Bakr, Soliman A. AUTHOR (S):

CORPORATE SOURCE: Bioanalytical and Drug Development Laboratory,

Biological and Medical Research (MBC-03), King Faisal

Specialist Hospital and Research Centre, Riyadh,

11211, Saudi Arabia

SOURCE: Chirality (1997), 9(1), 10-12

CODEN: CHRLEP; ISSN: 0899-0042

Wiley-Liss PUBLISHER: DOCUMENT TYPE: Journal English LANGUAGE:

Entered STN: 05 Apr 1997 ED

A newly developed reversed phase cellulose tris(4-Me benzoate) known as AΒ Chiralcel OJ-R was used to investigate the chiral recognition and enantiomeric separation of eight racemic piperidine-2,6-dione compds.-namely, aminoglutethimide and its major metabolite acetylaminoglutethimide, glutethimide, cyclohexylamino-glutethimide, pyridoglutethimide, thalidomide, phenglutarimide, and 3-phenylacetyl-amino-2,6-piperidinedione (antineoplaston A-10). Chiral separation of these compds. was achieved under varying ratios of the mobile phase, except for phenglutarimide and 3-phenylacetylamino-2,6-piperidinedione, for which separation was unsuccessful. Possible chiral recognition mechanisms are also presented.

CC 64-3 (Pharmaceutical Analysis)

ACCESSION NUMBER:

50-35-1, Thalidomide 77-21-4, Glutethimide IT 125-84-8, Aminoglutethimide 1121-89-7, Piperidine-2,6-dione 1156-05-4, Phenglutarimide 841-67-8 17575-59-6 38473-28-8, Acetylaminoglutethimide 2614-06-4 17575-58-5 57288-03-6 **77658-84-5** 55511-44-9 83155-00-4 91531-30-5, 92788-10-8, Pyridoglutethimide 108929-54-0 Antineoplaston A10 119095-49-7 112924-18-2 115883-22-2 121742-46-9 137623-90-6 137623-92-8

RL: ANT (Analyte); ANST (Analytical study) (enantiomeric separation of piperidine-2,6-dione drugs by HPLC using chiralcel OJ-R column)

L131 ANSWER 13 OF 44 CAPLUS COPYRIGHT 2005 ACS on STN

1996:750266 CAPLUS

DOCUMENT NUMBER:

126:51049

TITLE:

Enantioseparation of 3-phenylacetylamino-2,6-

piperidinedione and related chiral compounds

AUTHOR(S): Tang, Yubing; Reepmeyer, John C.; Revelle, Larry K.; Wilson, Joe A.

CORPORATE SOURCE: Division Drug Analysis, Food & Drug Administration,

St. Louis, MO, USA

SOURCE:

Journal of Chromatography, A (1996), 752(1+2), 93-99

CODEN: JCRAEY; ISSN: 0021-9673

PUBLISHER: Elsevier
DOCUMENT TYPE: Journal
LANGUAGE: English
ED Entered STN: 21 Dec 1996

This paper reports HPLC methodol. for the first successful enantiomeric sepns. of 3-phenylacetylamino-2,6-piperidinedione (PAP), a naturally occurring peptide derivative used for inhibiting the growth of cancer tissues. The chiral separation of four related hydrolyzates is also described. A com. available tris-4-methylbenzoate cellulose (Chiralcel OJ) column was used as the chiral stationary phase, operated in the normal-phase mode. The results demonstrated that hydrolyzed products of PAP, each of which has a carboxylic acid functionality present in its structure, eluted in a reasonable time and are enantiomerically resolved only when a trace amount of organic acid is present in the mobile phase. Different alcs. (ethanol and isopropanol) and acid additives (trifluoroacetic acid, trichloroacetic acid and acetic acid) were evaluated. In general, for the separation of the acidic enantiomers, ethanol is superior to isopropanol and stronger acids enhance the resolution more effectively. However, chiral separation of PAP could

CC 64-3 (Pharmaceutical Analysis)

additive was needed.

IT 2752-33-2 2752-34-3 2752-35-4 3343-29-1 7607-72-9 28047-15-6 **77658-84-5**, 3-Phenylacetylamino-2,6-piperidinedione 91531-30-5 104771-88-2 108929-54-0 174591-46-9 184849-21-6 184849-22-7

only be accomplished with isopropanol in the mobile phase and no acidic

184972-99-4 185019-08-3

RL: ANT (Analyte); ANST (Analytical study)

(enantiomeric separation of 3-phenylacetylamino-2,6-piperidinedione and related chiral compds. by HPLC)

L131 ANSWER 14 OF 44 CAPLUS COPYRIGHT 2005 ACS on STN

ACCESSION NUMBER: 1995:758941 CAPLUS

DOCUMENT NUMBER:

123:160816

TITLE:

Design of drugs involving receptor-ligand-DNA

interactions

INVENTOR(S):

Hendry, Lawrence B.

PATENT ASSIGNEE(S):

USA

SOURCE:

PCT Int. Appl., 63 pp.

CODEN: PIXXD2

DOCUMENT TYPE:

Patent

LANGUAGE:

English

FAMILY ACC. NUM. COUNT: 2

PATENT INFORMATION:

PATENT	NO.			KIN	D -	DATE		j	APPL	ICAT	ION 1	NO.		Di -	ATE	
WO 9514	791			A 1		1995	0601	1	WO 1:	994 -1	US13	765		1	9941	128
₩:	GB,	GE,	HU,	JP,	KE,	BR, KG, PL,	KP,	KR,	KZ,	LK,	LR,	LT,	LU,	LV,	MD,	MG,
RW:	•	MW, NL,			-	BE, BJ,	•		-		-					

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AU 1995-12979
    AU 9512979
                         A1
                               19950613
                                                                 19941128
                                          EP 1995-904188
    EP 740708
                               19961106
                                                                 19941128
                         Α1
    EP 740708
                               20040804
                         B1
        R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IE, IT, LI, LU, MC, NL, PT, SE
                                         JP 1994-515280
     JP 09505603
                        T2 19970603
                                                                19941128
                                           AT 1995-904188
    AT 272719
                               20040815
                                                                 19941128
                         Ε
                                           US 1993-158689
PRIORITY APPLN. INFO.:
                                                              A 19931126
                                           WO 1994-US13765
                                                             W 19941128
     Entered STN: 26 Aug 1995
     It has been discovered that the degree of hormonal activity of candidate
AΒ
     ligands correlates better with degree of fit into DNA than with the
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AB It has been discovered that the degree of hormonal activity of candidate ligands correlates better with degree of fit into DNA than with the strength of receptor binding, and that the receptors in the steroid/thyroid hormone/vitamin A and D family alter the physiochem. properties of DNA and in concert with other transcription factors facilitate insertion of the ligand into DNA. As a result, the magnitude of the response is a function of the structure of the ligand as it relates to insertion and fit into the DNA, and the specificity of the response is a function of the stereochem. of the receptor through binding to both the ligand and to the DNA. Based on these discoveries, a method is described herein for identifying drugs having increased activity as compared with the natural ligand for receptors, e.g. estrogenic receptors.

IC ICM C12Q001-68

ICS C07H021-02; C07H021-04

CC 1-3 (Pharmacology)

Section cross-reference(s): 2

77658-84-5, 3-Phenylacetylamino-2,6-piperidinedione 138592-72-0 167172-99-8, SGI 100 167173-00-4, SGI 101 167173-01-5, SGI 102 RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); PRP (Properties); BIOL (Biological study)

(drug design involving receptor-ligand-DNA interactions)

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L131 ANSWER 15 OF 44 CAPLUS COPYRIGHT 2005 ACS on STN
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ACCESSION NUMBER: 1994:680507 CAPLUS

DOCUMENT NUMBER: 121:280507

TITLE: Chemical modification of antineoplaston AlO and

antitumor activity of its analogs

AUTHOR(S): Huang, Junqin; Ma, Weiyong; Zhang, Chunnian

CORPORATE SOURCE: Shanghai Inst. Pharm. Ind., Shanghai, 200040, Peop.

Rep. China

SOURCE: Zhongguo Yiyao Gongye Zazhi (1993), 24(10), 437-41,

451

CODEN: ZYGZEA; ISSN: 1001-8255

DOCUMENT TYPE: Journal LANGUAGE: Chinese

ED Entered STN: 10 Dec 1994

AB Antineoplaston A10 analogs I (R = benzyl, substituted benzyl, naphthylmethyl, thenyl, bromothenyl, PhCH:CH, etc.) were prepared by N-acylation of 3-aminopiperidine-2,6-dione with RCO2H. I showed little or no activity at 100µg/mL against L1210 leukemia cell.

CC 27-16 (Heterocyclic Compounds (One Hetero Atom))

Section cross-reference(s): 1

91531-30-5DP, Antineoplaston A10, IT**77658-84-5P** 91393-02-1P 91531-30**-**5P 138592-85-5P 138592-87-7P 138592-89-9P analogs 158828-54-7P 158828-55**-**8P 158828-56-9P 158828-57-0P 158828-58-1P 158828-59**-**2P 158828-60-5P 158828-61-6P 158828-63-8P 158828-64-9P 158828-66**-**1P 158828-67-2P 158828-68-3P 158828-69-4P 158828-65-0P 158930-24-6P 194712-35-1P 158828-70-7P

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); SPN (Synthetic preparation); BIOL (Biological study); PREP (Preparation)

(preparation and antitumor activity of)

L131 ANSWER 16 OF 44 CAPLUS COPYRIGHT 2005 ACS on STN ACCESSION NUMBER: 1994:124340 CAPLUS

DOCUMENT NUMBER: 120:124340

TITLE: Antitumor activity of 3-phenylacetylamino-2,6-

piperidinedione and its computer modeled analogs

AUTHOR(S): Copland, John Alton, III

CORPORATE SOURCE: Med. Coll. Georgia, Augusta, GA, USA

SOURCE: (1992) 222 pp. Avail.: Univ. Microfilms Int., Order

No. DA9225145

From: Diss. Abstr. Int. B 1992, 53(6), 2632

DOCUMENT TYPE: Dissertation LANGUAGE: English

Entered STN: 19 Mar 1994

AB Unavailable

ED

CC 1-6 (Pharmacology)

IT 77658-84-5, 3-Phenylacetylamino-2,6-piperidinedione

77658-84-5D, 3-Phenylacetylamino-2,6-piperidinedione, analogs RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL

(Biological study); USES (Uses) (antitumor activity of)

L131 ANSWER 17 OF 44 CAPLUS COPYRIGHT 2005 ACS on STN

ACCESSION NUMBER: 1994:124306 CAPLUS

DOCUMENT NUMBER: 120:124306

TITLE: In vitro and in vivo studies on the antineoplastic

properties and mechanism of action of a novel amino acid analog, 3-phenylacetylamino-2,6-piperidinedione

(A10)

AUTHOR(S): Wood, Joseph Clifton

CORPORATE SOURCE: Med. Coll. Georgia, Augusta, GA, USA

SOURCE: (1992) 148 pp. Avail.: Univ. Microfilms Int., Order

No. DA9225153

From: Diss. Abstr. Int. B 1992, 53(5), 2201

DOCUMENT TYPE: Dissertation

LANGUAGE: English
ED Entered STN: 19 Mar 1994

AB Unavailable

CC 1-6 (Pharmacology)

IT 77658-84-5, 3-Phenylacetylamino-2,6-piperidinedione

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL

(Biological study); USES (Uses)

(antitumor activity of, mechanism of)

L131 ANSWER 18 OF 44 CAPLUS COPYRIGHT 2005 ACS on STN

ACCESSION NUMBER: 1992:188060 CAPLUS

DOCUMENT NUMBER: 116:188060

TITLE: Substituted piperidinediones, substituted phenylacetic

acids, and substituted phenylacetylglutamines for

treatment of AIDS

INVENTOR(S): Burzynski, Stanislaw R.

PATENT ASSIGNEE(S): USA

SOURCE: U.S., 5 pp.

CODEN: USXXAM

DOCUMENT TYPE: Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO. KIND DATE APPLICATION NO. DATE

```
US 1990-577464
    US 5089508
                         Α
                               19920218
                                                                  19900904
                                           WO 1991-US5977
    WO 9204027
                         A1
                               19920319
                                                                 19910821
        W: AT, AU, BB, BG, BR, CA, CH, CS, DE, DK, ES, FI, GB, HU, JP, KP,
            KR, LK, LU, MC, MG, MN, MW, NL, NO, PL, RO, SD, SE, SU
        RW: AT, BE, BF, BJ, CF, CG, CH, CI, CM, DE, DK, ES, FR, GA, GB, GN,
            GR, IT, LU, ML, MR, NL, SE, SN, TD, TG
                               19920330
                                          AU 1991-85397
                                                                  19910821
    AU 9185397
                         Α1
    AU 638869
                         B2
                               19930708
    EP 500905
                               19920902
                                          EP 1991-917237
                                                                  19910821
                         Α1
    EP 500905
                         В1
                               19960313
        R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE
                               19960315 AT 1991-917237
    AT 135217
                        E
                                                                19910821
                                          ES 1991-917237
    ES 2084181
                        Т3
                               19960501
                                                                  19910821
    SG 81889
                        A1
                                          SG 1996-7278
                               20010724
                                                                  19910821
                        Α
                                          ZA 1991-6977
    ZA 9106977
                               19920624
                                                                 19910903
                        Α
                                          US 1991-790584
    US 5254587
                               19931019
                                                                 19911108
                                         US 1992-888976
    US 5244922
                        Α
                               19930914
                                                                 19920527
                                        WO 1993-US5002
    WO 9324123
                         A1
                               19931209
                                                                  19930526
            AT, AU, BB, BG, BR, CA, CH, CZ, DE, DK, ES, FI, GB, HU, JP, KP,
            KR, LK, LU, MG, MN, MW, NL, NO, NZ, PL, PT, RO, RU, SD, SE, SK,
            UA, VN
         RW: AT, BE, CH, DE, DK, ES, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE,
            BF, BJ, CF, CG, CI, CM, GA, GN, ML, MR, NE, SN, TD, TG
                               19931230 AU 1993-43927
     AU 9343927
                         Α1
                                                                 19930526
                                          EP 1993-914168
                               19940615
    EP 601164
                         Α1
                                                                  19930526
                               20000119
    EP 601164
                         В1
        R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IE, IT, LI, LU, MC, NL, PT, SE
                                        AT 1993-914168
                               20000215
     AT 188871
                        Ε
                                                                19930526
                                           ES 1993-914168
                         Т3
                                                                  19930526
    ES 2142873
                               20000501
                         A1
                                           HK 1998-115815
                                                                  19981228
    HK 1014499
                               20000804
PRIORITY APPLN. INFO.:
                                           US 1990-577464
                                                             A 19900904
                                                             A 19910821
                                           WO 1991-US5977
                                                             A2 19911108
                                           US 1991-790584
                                                             A 19920527
                                           US 1992-888976
                                                             A 19930526
                                           WO 1993-US5002
OTHER SOURCE(S):
                        MARPAT 116:188060
    Entered STN: 16 May 1992
ED
    Piperidinedione derivs. I [R = H, OH, NH2, OW; W = C1-12 aliphatic, C(O)Z; Z
AB
     = C1-12 aliphatic or aromatic; X = H, F, Cl, Br, I, OH, OW (W as above), NO2,
    NH2; Y = H, F, Cl, Br, I], and their pharmaceutically acceptable salts,
     are claimed for treatment of AIDS. Also disclosed for treatment of AIDS
     are hydrolysis products of I, i.e., substituted phenylacetic acids and
     substituted phenylacetylglutamines. Efficacy of antineoplaston AS2-1 (1:4
     ratio of Na salt of phenylacetylglutamine and Na salt of phenylacetic
     acid) in clin. case studies is reported.
     ICM A61K031-445
IC
NCL
    514328000
     1-5 (Pharmacology)
CC
    77658-84-5 77658-84-5D, derivs. 104624-98-8,
ΙT
     Antineoplaston AS2-1
    RL: BIOL (Biological study)
        (AIDS treatment with)
L131 ANSWER 19 OF 44 CAPLUS COPYRIGHT 2005 ACS on STN
ACCESSION NUMBER:
                      1990:497459 CAPLUS
DOCUMENT NUMBER:
                        113:97459
TITLE:
                        Preparation of 3-[(phenylacetyl)amino]piperidine-2,6-
                        dione and its use as antineoplastic agents
INVENTOR(S):
                        Burzynski, Stanislaw R.
PATENT ASSIGNEE(S):
                        USA
                        U.S., 5 pp.
SOURCE:
```

CODEN: USXXAM

DOCUMENT TYPE: Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
US 4918193	Α	19900417	US 1989-295372	19890111
FI 9000129	Α	19900712	FI 1990-129	19900110
FI 92391	В	19940729		
FI 92391	С	19941110		
SU 1809830	A3	19930415	SU 1990-4742866	19900110
PL 163552	B1	19940429	PL 1990-283254	19900110
KR 139204	B1	19980515	KR 1990-205	19900110
LT 3518	В	19951127	LT 1993-681	19930623
PRIORITY APPLN. INFO.:			US 1989-295372	A 19890111

ED Entered STN: 16 Sep 1990

The title compound (I) was prepared in greater yield by an improved procedure comprising mixing L-glutamine and a phenylacetyl halide, preferably chloride, in a weakly alkaline aqueous solution, adjusting the pH to 2-3, and cyclization of the intermediate phenylacetyl glutamine by heating the reaction mixture to .apprx.160°. I-Na salt upon standing undergoes basic hydrolysis to form Na salts of its degradation products: phenylacetyl glutamine (II) and phenylacetylisoglutamine (III). In a bioassay in vitro 2 mg I/mL, 10 mg II/mL, and 3 mg/mL of a 1:4 mixture of III with C6H5CH2CO2H stabilized the number of MDA-MB-231 human breast carcinoma cells counted after 24 h from incubation and persisting for ≥48 h. The results of phase I clin. trials with I, II, III, and C6H5CH2CO2H and their mixts., involving various neoplastic diseases, were reported. The prepns. of I for injection, capsule and solns. were also given.

IC ICM C07D211-40

NCL 546220000

CC 27-16 (Heterocyclic Compounds (One Hetero Atom))

Section cross-reference(s): 1, 63

TT 77658-84-5P, 3-[(Phenylacetyl)amino]piperidine-2,6-dione RL: SPN (Synthetic preparation); PREP (Preparation)

(preparation of, as antineoplastic agent)

L131 ANSWER 20 OF 44 CAPLUS COPYRIGHT 2005 ACS on STN

ACCESSION NUMBER: 1988:49339 CAPLUS

DOCUMENT NUMBER: 108:49339

TITLE: Use of 3-N-phenylacetylamino-2,6-piperidinedione for

treatment of neuropsychiatric disorders

INVENTOR(S): Hendry, Lawrence B.; Diamond, Ana H.; Diamond, Bruce

I.; Ewing, Douglas E.

PATENT ASSIGNEE(S): Stereochemical Genetics, Inc., USA

SOURCE: U.S., 5 pp. CODEN: USXXAM

DOCUMENT TYPE: Patent

LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE	
US 4705796	Α	19871110	US 1986-899822	19860825	
PRIORITY APPLN. INFO.:			US 1986-899822	19860825	

ED Entered STN: 20 Feb 1988

AB Disorders involving monoamine oxidase(MAO) regulator are treated with 3-N-phenylacetylamnino-2,6-piperidinedione(I), which is an effective and selective MAO type B inhibitor. The effect of I on platelet MAO B

09/955010 Jones Page 22

activity was studied in vitro from blood collected from normal volunteers. Compared with pargyline, I was a less potent MAO B inhibitor. In human and rat brain in vitro, I preferentially inhibited the MAO type B enzyme at concns. 10 time less than needed to affect MAO A activity.

IC ICM A61K031-445

NCL 514328000

1-11 (Pharmacology) CC

77658-84-5 TΤ

RL: BIOL (Biological study)

(pharmaceutical containing, for neuropsychiatric disorder treatment)

L131 ANSWER 21 OF 44 CAPLUS COPYRIGHT 2005 ACS on STN

ACCESSION NUMBER: 1984:635560 CAPLUS

DOCUMENT NUMBER:

101:235560

TITLE:

Purified antineoplaston fractions and methods of

treating neoplastic disease

INVENTOR(S):

Burzynski, Stanislaw R.

PATENT ASSIGNEE(S):

USA

U.S., 14 pp. Cont.-in-part of U.S. Ser. No. 279,728, SOURCE:

abandoned. CODEN: USXXAM

DOCUMENT TYPE:

Patent

LANGUAGE:

English

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

PATENT NO.		DATE	APPLICATION NO.	DATE
US 4470970		19840911	US 1981-330383	19811215
CA 1188218	A1	19850604	CA 1982-403789	19820526
AU 8284239	A1	19830106	AU 1982-84239	19820527
AU 551109	B2	19860417		
DK 8202420	Α	19830103	DK 1982-2420	19820528
DK 162813		19911216		
DK 162813				
EP 69232	A2	19830112	EP 1982-104867	19820603
EP 69232	A 3	19840704	·	
EP 69232	B1	19861029		
R: AT, BE, CH				
IL 65960	A1	19851129	IL 1982-65960	19820603
AT 23113	E	19861115	AT 1982-104867	19820603
ES 512894	A 1	19850101	ES 1982-512894	
ZA 8204178	Α	19830629 19830103	ZA 1982-4178	19820614
NO 8202218	Α	19830103	NO 1982-2218	19820629
NO 163595	В С	19900319		
NO 163595	C	19900627		
JP 58010521	A2	19830121	JP 1982-115330	19820702
JP 07029925	B4	19950405		
US 4558057	Α	19851210	US 1984-642499	19840820
US 4559325	Α	19851217	US 1984-642873	
CA 1262907	A2	19891114	CA 1985-475098	19850225
JP 05032548	A2	19930209	JP 1991-133639	19910509
JP 07039390	B4	19950501		
JP 05058886		19930309	JP 1991-133638	19910509
JP 07080764	B4	19950830		
DK 9101434	Α	19910806	DK 1991-1434	19910806
PRIORITY APPLN. INFO.:			US 1981-279728	A2 19810702
			US 1981-330383	A 19811215
			CA 1982-403789	
			EP 1982-104867	A 19820603

ED Entered STN: 22 Dec 1984

Highly purified fractions from human urine exhibit antineoplastic AB

Page 23

activity. The comprise biol. active small sized, low mol. weight peptides. 3-(Phenylacetylamino)piperidine-2,6-dione (I) [77658-84-5] was isolated from these fractions and showed antineoplastic activity. I was synthesized from these latter 2 compds. I was hydrolyzed 1st to N-phenylacetylglutamine [28047-15-6] and then further to PhCH2CO2H [103-82-2] and glutamine [56-85-9]. Parenteral solns. were prepared by reconstituting antineoplastic fractions, I, and its degradation products in pyrogen free H2O. The fractions showed antineoplastic activity against human breast carcinoma cultures and in patients with cancer, with I showing the highest activity.

IC A61K037-00; C07C103-52; C07G007-00

NCL 424177000

CC 63-3 (Pharmaceuticals)

Section cross-reference(s): 1, 2, 27

IT 77658-84-5

RL: BIOL (Biological study)

(of urine, as neoplasm inhibitor)

L131 ANSWER 22 OF 44 CAPLUS COPYRIGHT 2005 ACS on STN

ACCESSION NUMBER: 1983:137623 CAPLUS

DOCUMENT NUMBER: 98:137623

TITLE: Purified antineoplaston fractions and methods of

treating neoplastic disease

INVENTOR(S): Burzynski, Stanislaw R.

PATENT ASSIGNEE(S): USA

SOURCE: Eur. Pat. Appl., 39 pp.

CODEN: EPXXDW

DOCUMENT TYPE: Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT: 2

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.		DATE
				-	
EP 69232	A2	19830112	EP 1982-104867		19820603
EP 69232	A3	19840704			
EP 69232	B1	19861029	•		
R: AT, BE, CH,	DE, FR	, GB, IT, LI	I, LU, NL, SE		
US 4470970	Α	19840911	US 1981-330383		19811215
AT 23113	E	19861115	AT 1982-104867		19820603
PRIORITY APPLN. INFO.:			US 1981-279728	Α	19810702
			US 1981-330383	Α	19811215
			EP 1982-104867	Α	19820603

ED Entered STN: 12 May 1984

Antineoplastons (substances produced by a living organism which protect it against the development of neoplastic growth by a nonimmunol. process and which do not significantly inhibit the growth of normal tissues) were isolated from human urine by ultrafiltration (to eliminate compds. with mol. wts. >2000-5000), followed by diverse separation procedures. A common component of all the fractions was 3-(N-phenylacetylamino)piperidine-2,6-dione (I) [77658-84-5]. These fractions exhibited antineoplastic activity against cultures of human breast carcinoma and against various human cancers when used clin. Two hydrolysis degradation products of I, N-(phenylacetyl)glutamine [28047-15-6] and phenylacetic acid [103-82-2], also exhibited antineoplastic activity. I was synthesized by reacting a NaHCO3 solution of L-glutamine [56-85-9] with phenylacetyl chloride [103-80-0].

- IC A61K037-02; A61K035-22; C07C103-52
- CC 1-6 (Pharmacology)
- IT 77658-84-5P

RL: SPN (Synthetic preparation); PREP (Preparation) (preparation and extraction from human urine and antineoplastic activity of)

L131 ANSWER 23 OF 44 USPATFULL on STN

ACCESSION NUMBER:

2003:153436 USPATFULL

NUMBER

TITLE:

Formulation of amino acids and riboflavin useful to reduce toxic effects of cytotoxic chemotherapy

DATE

20030605

20011127

KIND

A1

Α1

Burzynski, Stanislaw R., Houston, TX, UNITED STATES INVENTOR (S):

PATENT INFORMATION: US 2003105104 APPLICATION INFO.: US 2001-995010 DOCUMENT TYPE: Utility FILE SEGMENT: APPLICATION

HOWREY SIMON ARNOLD & WHIT LEGAL REPRESENTATIVE:

TX, 77057-2198

NUMBER OF CLAIMS: EXEMPLARY CLAIM: LINE COUNT: 834

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

Pharmaceutical compositions effective in alleviating or reducing the effects of fatigue and weakness associated with cancer and cytotoxic cancer chemotherapy are disclosed. The pharmaceutical compositions of the present invention comprise ribo lavin, effectors of the urea cycle in free form or pharmacologically acceptable salts thereof, and amino acids selected from the groups of essential and non-essential amino acids, in free form or pharmaceutically acceptable salts thereof, suitably combined with appropriate carriers, diluents, or excipients. Also disclosed are methods of alleviating or reducing the effects of fatigue and weakness associated with cancer and cytotoxic cancer chemotherapy by administration of pharmaceutical compositions of the present invention.

56-40-6, Glycine, biological studies 56-41-7, Alanine, IT biological studies 56-45-1, Serine, biological studies 70-26-8, Ornithine 72-18-4, Valine, biological studies 72-19-5, Threonipe, biological studies 74-79-3, Arginine, biological studies 83-88-5, Riboflavin, biological studies 107-35-7, Taurine 372-75-8, Citrulline 77658-84-5, \$\mathscr{A}\$-Phenylacetylamino-2,6-piperidinedione

(formulation of amino acids and riboflavin useful to reduce toxic effects of cytotoxic chemotherapy)

L131 ANSWER 24 OF 44 USPATFULL on STN

ACCESSION NUMBER: 2002:126278 USPATFULL

TITLE: Design of drugs involving receptor-ligand-DNA

interactions

INVENTOR(S): Hendry, Lawrence B., Augusta, CA, UNITED STATES

NUMBER KIND DATE PATENT INFORMATION: US 2002064790 **A1** 20020530 APPLICATION INFO.: US 2001-941230 20010828 **A1** (9) Continuation of Ser. No. US 1999-239491, filed on 28 RELATED APPLN. INFO.:

Jan 1999, PATENTED Continuation of Ser. No. US 1997-864669, filed on 28 May 1997, PATENTED

Continuation of Ser. No. US 1994-369779, filed on 28 Nov 1994, PATENTED Continuation-in-part of Ser. No. US 1993-158689, filed on 26 Nov 1993, ABANDONED

DOCUMENT TYPE: Utility

APPLICATION FILE SEGMENT:

LEGAL REPRESENTATIVE: John S. Pratt, Esq., KILPATRICK STOCKTON LLP, Suite

2800, 1100 Peachtree Street, Atlanta, GA, 30309-4530

NUMBER OF CLAIMS: 26 EXEMPLARY CLAIM: 1

NUMBER OF DRAWINGS: 7 Drawing Page(s)

LINE COUNT: 1395

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

It has been discovered that the degree of hormonal activity of candidate ligands correlates better with degree of fit into DNA than with the strength of receptor binding, and that the receptors in the steroid/thyroid hormone/vitamin A and D family alter the physiochemical properties of DNA and in concert with other transcription factors facilitate insertion of the ligand into DNA. As a result, the magnitude of the response is a function of the structure of the ligand as it related to insertion and fit into the DNA and the specificity of the response is a function of the stereochemistry of the receptor through binding to both the ligand and to the DNA. Based on these discoveries, a method is described herein for identifying drugs having increased activity as compared with the natural ligand for receptors such as the estrogenic receptors.

L131 ANSWER 25 OF 44 USPATFULL on STN

ACCESSION NUMBER: 2001:185035 USPATFULL

TITLE: Design of drugs involving receptor-ligand-DNA

interactions

INVENTOR(S): Hendry, Lawrence B., 1939 Bolin Rd., North Augusta, SC,

United States 29841

RELATED APPLN. INFO.: Continuation of Ser. No. US 1997-864669, filed on 28

May 1997, now patented, Pat. No. US 5888738

Continuation of Ser. No. US 1994-369779, filed on 28

Nov 1994, now patented, Pat. No. US 5705335

Continuation-in-part of Ser. No. US 1993-158689, filed

on 26 Nov 1993, now abandoned

DOCUMENT TYPE: Utility FILE SEGMENT: GRANTED

PRIMARY EXAMINER: Brusca, John S.

LEGAL REPRESENTATIVE: Kilpatrick Stockton LLP

NUMBER OF CLAIMS: 6 EXEMPLARY CLAIM: 1

NUMBER OF DRAWINGS: 9 Drawing Figure(s); 9 Drawing Page(s)

LINE COUNT: 1345

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

AB It has been discovered that the degree of hormonal activity of candidate ligands correlates better with degree of fit into DNA than with the strength of receptor binding, and that the receptors in the steroid/thyroid hormone/vitamin A and D family alter the physiochemical properties of DNA and in concert with other transcription factors facilitate insertion of the ligand into DNA. As a result, the magnitude of the response is a function of the structure of the ligand as it related to insertion and fit into the DNA and the specificity of the response is a function of the stereochemistry of the receptor through binding to both the ligand and to the DNA. Based on these discoveries, a method is described herein for identifying drugs having increased activity as compared with the natural ligand for receptors such as the estrogenic receptors.

L131 ANSWER 26 OF 44 USPATFULL on STN

ACCESSION NUMBER: 1999:40164 USPATFULL

TITLE: Computer-based design and screening of molecules using

DNA interactions

INVENTOR(S): Hendry, Lawrence B., 1939 Bolin Rd., North Augusta, SC,

United States 29841

RELATED APPLN. INFO.: Division of Ser. No. US 1994-369779, filed on 28 Nov

1994, now patented, Pat. No. US 5705335 which is a continuation-in-part of Ser. No. US 1993-158689, filed

on 26 Nov 1993, now abandoned

DOCUMENT TYPE: Utility
FILE SEGMENT: Granted
PRIMARY EXAMINER: Degen, Nancy
ASSISTANT EXAMINER: Brusca, John S.
LEGAL REPRESENTATIVE: Jones & Askew, LLP

NUMBER OF CLAIMS: 31 EXEMPLARY CLAIM: 1

NUMBER OF DRAWINGS: 22 Drawing Figure(s); 9 Drawing Page(s)

LINE COUNT: 1453

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

It has been discovered that the degree of hormonal activity of candidate ligands correlates better with degree of fit into DNA than with the strength of receptor binding, and that the receptors in the steroid/thyroid hormone/vitamin A and D family alter the physiochemical properties of DNA and in concert with other transcription factors facilitate insertion of the ligand into DNA. As a result, the magnitude of the response is a function of the structure of the ligand as it related to insertion and fit into the DNA and the specificity of the response is a function of the stereochemistry of the receptor through binding to both the ligand and to the DNA. Based on these discoveries, a method is described herein for identifying drugs having increased activity as compared with the natural ligand for receptors such as the estrogenic receptors.

L131 ANSWER 27 OF 44 USPATFULL on STN

ACCESSION NUMBER: 1999:40161 USPATFULL

TITLE: Design of drugs involving receptor-ligand-DNA

interactions

INVENTOR(S): Hendry, Lawrence B., 1939 Bolin Rd., North Augusta, SC,

United States 29841

RELATED APPLN. INFO.: Continuation of Ser. No. US 1994-369779, filed on 28

Nov 1994, now patented, Pat. No. US 5705335 which is a continuation-in-part of Ser. No. US 1993-158689, filed

on 26 Nov 1993, now abandoned

DOCUMENT TYPE: Utility
FILE SEGMENT: Granted
PRIMARY EXAMINER: Degen, Nancy

ASSISTANT EXAMINER: Brusca, John S.
LEGAL REPRESENTATIVE: Jones & Askew, LLP

NUMBER OF CLAIMS: 23 EXEMPLARY CLAIM: 1

NUMBER OF DRAWINGS: 9 Drawing Figure(s); 9 Drawing Page(s)

LINE COUNT: 1407

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

It has been discovered that the degree of hormonal activity of candidate ligands correlates better with degree of fit into DNA than with the strength of receptor binding, and that the receptors in the steroid/thyroid hormone/vitamin A and D family alter the physiochemical properties of DNA and in concert with other transcription factors facilitate insertion of the ligand into DNA. As a result, the magnitude of the response is a function of the structure of the ligand as it related to insertion and fit into the DNA and the specificity of the response is a function of the stereochemistry of the receptor through binding to both the ligand and to the DNA. Based on these discoveries, a method is described herein for identifying drugs having increased activity as compared with the natural ligand for receptors such as the estrogenic receptors.

IT 77658-84-5, 3-Phenylacetylamino-2,6-piperidinedione (drug design involving receptor-ligand-DNA interactions)

L131 ANSWER 28 OF 44 IPA COPYRIGHT 2005 ASHP on STN DUPLICATE 2

ACCESSION NUMBER: 2003:11775 IPA

DOCUMENT NUMBER: 40-11775

TITLE: Phase II study of antineoplaston A10 and AS2-1 in patients

with recurrent diffuse intrinsic brain stem glioma - A

preliminary report

AUTHOR: Burzynski, SR; Lewy, RI; Weaver, RA; Axler, ML; Bestak, M;

et al

CORPORATE SOURCE: Burzynski Clin, Dept Internal Med, 9432 Old Katy Rd,

Houston, TX, USA info@burzynskiclinic.com

SOURCE: Drugs in R and D, (2003) Vol. 4, pp. 91-101. 41 Refs.

CODEN: DRURDA; ISSN: 1174-5886.

DOCUMENT TYPE: Journal FILE SEGMENT: HUMAN LANGUAGE: English

ABSTRACT:

Objective: A phase 2 study of antineoplaston A10 and AS2-1 was conducted to evaluate the antineoplastic activity in patients with recurrent diffuse intrinsic brain stem glioma.

Patients and methods: This report describes the results of treatment of the first 12 patients admitted to the study. Patients received escalating doses of antineoplaston A10 and AS2-1 by intravenous bolus injections. The median duration of treatment was 6 months and the average dosage of antineoplaston A10 was 11.3 g/kg/day and of antineoplaston AS2-1 0.4 g/kg/day. Responses were assessed by gadolinium-enhanced magnetic resonance imaging of the head.

Results: Of ten evaluable patients, complete response was determined in two cases (20%), partial response in three (30%), stable disease in three (30%) and progressive disease in two (20%). Survival at 2 years was 33.3%. Currently, of all 12 patients, two (17%) were alive and tumour free for over 5 years since initial diagnosis; one was alive for more than 5 years, and another for more than 4 years from the start of treatment. Only mild and moderate toxicities were observed, which included three cases of skin allergy, two cases of anaemia, fever and hypernatraemia, and single cases of agranulocytosis, hypoglycaemia, numbness, tiredness, myalgia and vomiting.

Conclusion: The results of this study compared favourably with the

responses of patients treated with radiation therapy and chemotherapy

. The study continues with accrual of additional patients.

SECTION: 5 Investigational Drugs; 4 Toxicity

CLASSIFICATION: 10:00 Antineoplastic agents; 10:00 Antineoplastic agents

INDEX TERM: Antineoplaston A10; glioma
INDEX TERM: Antineoplaston AS2-1; glioma
INDEX TERM: Glioma; antineoplaston A10
INDEX TERM: Dosage; antineoplaston A10
INDEX TERM: Toxicity; antineoplaston A10

INDEX TERM: Antineoplastic agents; antineoplaston A10

INDEX TERM: Glioma; antineoplaston AS2-1
INDEX TERM: Dosage; antineoplaston AS2-1
INDEX TERM: Toxicity; antineoplaston AS2-1

INDEX TERM: Antineoplastic agents; antineoplaston AS2-1

CAS REGISTRY NO.: 77658-84-5 (Antineoplaston A10)
CAS REGISTRY NO.: 104624-98-8 (Antineoplaston AS2-1)

L131 ANSWER 29 OF 44 IPA COPYRIGHT 2005 ASHP on STN DUPLICATE 3

ACCESSION NUMBER: 2000:11825 IPA

DOCUMENT NUMBER: 37-11826

TITLE: Retrospective study of antineoplastons A10 and AS2-1 in

primary brain tumors

AUTHOR: Burzynski, S. R.; Conde, A. B.; Peters, A.; Saling, B.;

Nacht, C. H.; et al

CORPORATE SOURCE: Burzynski Clin., 12000 Richmond Ave., Houston, TX

77082-2431, USA Internet: jpaszkowia@aol.com

SOURCE: Clinical Drug Investigation (New Zealand), (Jul 1999) Vol.

18, pp. 1-10. 30 Refs.

CODEN: CDINFR; ISSN: 1173-2563.

DOCUMENT TYPE: Journal FILE SEGMENT: HUMAN

LANGUAGE: English

ABSTRACT:

To evaluate the new treatment of brain tumors with antineoplastons, a study of 36 patients (median age 38.5 yr) who had failed established therapies and received daily intravenous injections of antineoplastons A10 and AS2-1 at average dosages of 7.7 and 0.36 g/kg/day, respectively, was conducted.

Antineoplaston therapy eliminated or substantially reduced tumors in 44% of patients with brain tumors. It produced complete response in 9 patients, partial response in 7 patients, stable disease in 12 patients, and progressive disease in 8 patients. Adverse drug effects included skin rash, somnolence, weakness, nausea, vomiting, slurred speech, and abnormalities in plasma electrolytes, and were reversed on temporary discontinuation or dose reduction. Compared with standard treatment, antineoplaston therapy was associated with prolonged survival time and prolonged time to disease progression.

Yinghua Shu Wang

SECTION: 5 Investigational Drugs; 4 Toxicity

CLASSIFICATION: 10:00 Antineoplastic agents; 10:00 Antineoplastic agents

INDEX TERM: Antineoplaston A10; brain neoplasms
INDEX TERM: Antineoplaston AS2-1; brain neoplasms
INDEX TERM: Brain neoplasms; antineoplaston A10
INDEX TERM: Brain neoplasms; antineoplaston AS2-1

INDEX TERM: Antineoplastic agents; antineoplaston A10; brain neoplasms

INDEX TERM: Antineoplastic agents; antineoplaston AS2-1; brain

neoplasms

INDEX TERM: Toxicity; antineoplaston Al0
INDEX TERM: Toxicity; antineoplaston AS2-1
CAS REGISTRY NO.: 77658-84-5 (Antineoplaston AS2-1)
CAS REGISTRY NO.: 104624-98-8 (Antineoplaston AS2-1)

09/955010 Jones Page 29

L131 ANSWER 30 OF 44 IPA COPYRIGHT 2005 ASHP on STN DUPLICATE 7

86:4535 IPA ACCESSION NUMBER: 24-05051 DOCUMENT NUMBER:

Toxicology studies on antineoplaston A10 injections in TITLE:

cancer patients

Burzynski, S. R.; Kubove, E. **AUTHOR:**

CORPORATE SOURCE: Burzynski Res. Inst., Inc., 12707 Trinity Drive, Stafford,

TX 77477

Drugs Under Experimental and Clinical Research SOURCE:

(Switzerland), (1986) Vol. 12, pp. 47-55. 11 Refs.

CODEN: DECRDR; ISSN: 0378-6501.

DOCUMENT TYPE: Journal FILE SEGMENT: **MAMUH** LANGUAGE: English

ABSTRACT:

Toxicology studies of antineoplaston A10 (3-phenylacetylamino-2,6piperidinedione) in 18 cancer patients following intravenous injection of up to 2.21 g/kg/day for 52 to 640 days are described.

The treatment was associated with minimal side effects including febrile reactions, muscle and joint pain, muscle contraction in the throat, abdominal pain, nausea, dizziness and headache. Objective response was noticed in 8 patients.

Victor Origoni

SECTION: 4 Toxicity; 5 Investigational Drugs

CLASSIFICATION: 10:00 Antineoplastic agents INDEX TERM: Antineoplaston A10; toxicity

Toxicity; antineoplaston A10; side INDEX TERM:

effects

INDEX TERM: Antineoplastic agents; antineoplaston A10; toxicity

CAS REGISTRY NO.: 77658-84-5 (Antineoplaston A10)

CHEMICAL NAME: Antineoplaston A10 (3-Phenylacetylamino-2,6-

piperidinedione)

L131 ANSWER 31 OF 44 IPA COPYRIGHT 2005 ASHP on STN DUPLICATE 8

ACCESSION NUMBER: 84:10760 IPA 23-06501 DOCUMENT NUMBER:

Human toxicology studies on oral formulation of TITLE:

antineoplaston A10

Burzynski, S. R.; Mohabbat, M. O.; Burzynski, B. AUTHOR:

CORPORATE SOURCE: Burzynski Res. Inst., 12707 Trinity Dr., Stafford, TX 77477

Drugs Under Experimental and Clinical Research SOURCE:

(Switzerland), (1984) Vol. 10, pp. 891-909. 7 Refs. CODEN: DECRDR; ISSN: 0378-6501.

DOCUMENT TYPE: Journal FILE SEGMENT: HUMAN LANGUAGE: English

ABSTRACT:

The long term effect of 0.5 g oral antineoplaston A10 (I) capsule was evaluated in 42 patients (aged 17-72 yr) with 49 types of advanced neoplastic disease given 3-4 I capsules every 3-6 h.

The treatment with I was associated with minimal adverse reactions which included excessive gas in the stomach, gastrointestinal bleeding (probably unrelated to I), maculopapular rash, moderately increased blood pressure, vertigo, hypoglycemia and mild myelosuppression. At least some positive response manifested by clinical improvement was found in 75% of the 49 treated cases. The additional beneficial effects of I treatment included a decrease in plasma levels of triglycerides and cholesterol, an increase in white blood cell count and platelet count and the improvement of blood clotting.

It was concluded that more extensive clinical testing will be necessary to establish the effectiveness of I in neoplasm therapy.

Nancy F. Cruz

SECTION: 4 Toxicity; 5 Investigational Drugs

CLASSIFICATION: 10:00 Antineoplastic agents

INDEX TERM: Antineoplaston A10; oral; long term effects, neoplasms
INDEX TERM: Antineoplastic agents; antineoplaston A10; oral, long term

effects, neoplasms

INDEX TERM: Dosage schedules; antineoplaston A10; long term, effects INDEX TERM: Neoplasms; antineoplaston A10; oral, long term effects

INDEX TERM: Toxicity; antineoplaston A10; side

effects

CAS REGISTRY NO.: 77658-84-5 (Antineoplaston A10)

L131 ANSWER 32 OF 44 IPA COPYRIGHT 2005 ASHP on STN DUPLICATE 9

ACCESSION NUMBER: 84:10035 IPA DOCUMENT NUMBER: 23-02080

TITLE: Toxicology studies on oral formulation of antineoplaston

Alo in cancer patients

AUTHOR: Burzynski, S. R.; Mohabbat, M. O.; Burzynski, B.

CORPORATE SOURCE: Burzynski Res. Inst., Inc., 12707 Trinity Dr., Stafford, TX

77477

SOURCE: Drugs Under Experimental and Clinical Research

(Switzerland), (1984) Vol. 10, pp. 611-619. 8 Refs.

CODEN: DECRDR; ISSN: 0378-6501.

DOCUMENT TYPE: Journal FILE SEGMENT: HUMAN LANGUAGE: English

ABSTRACT:

The effect of chronic administration of antineoplaston A10 (3-phenylacetylamino-2,6-piperidinedione) was studied in 42 patients with advanced neoplastic diseases: the patients received 500 mg capsules for periods of 6 to 314 days.

The highest dose administered was 14 g over 24 h. Most patients were treated for 50 to 149 days. Therapy was associated with mild **side** ***effects*** , these included excess gas in the stomach, GI bleeding, maculopapular rash, moderately increased blood pressure, vertigo, hypoglycemia, hypokalemia and mild myelosuppression.

D. L. Thompson

SECTION: 5 Investigational Drugs
CLASSIFICATION: 10:00 Antineoplastic agents
INDEX TERM: Antineoplaston A10; toxicity

INDEX TERM: Antineoplastic agents; antineoplaston A10; toxicity

INDEX TERM: Toxicity; antineoplaston A10
CAS REGISTRY NO.: 77658-84-5 (Antineoplaston A10)

CHEMICAL NAME: Antineoplaston A10 (3-Phenylacetylamino-2,6-

piperidinedione)

L131 ANSWER 33 OF 44 BIOSIS COPYRIGHT (c) 2005 The Thomson Corporation. on

STN

ACCESSION NUMBER: 2001:99118 BIOSIS DOCUMENT NUMBER: PREV200100099118

TITLE: Novel piperidinedione analogs as inhibitors of breast

cancer cell growth.

AUTHOR(S): Abou-Zeid, L. A.; El-Mowafy, A. M. [Reprint author];

El-Ashmawy, M. B.; Hendry, L. B.; Abdelal, A. M.; Badria,

F. A.

CORPORATE SOURCE: Department of Applied Therapeutics, Faculty of Pharmacy,

Kuwait University, Safat, Kuwait

SOURCE: Archiv der Pharmazie (Weinheim), (December, 2000) Vol. 333,

No. 12, pp. 431-434. print. CODEN: ARPMAS. ISSN: 0365-6233.

DOCUMENT TYPE: Article

LANGUAGE:

English

ENTRY DATE:

Entered STN: 21 Feb 2001

Last Updated on STN: 15 Feb 2002

ABSTRACT:We previously reported the utility of antineoplaston-A10 (3-phenylacetylamino-2,6-piperidinedione) as an endogenous cancer protector and immune modulator in breast cancer patients (Cancer Lett., 2000, 157, 57). In this study, four new piperidinedione A10 analogs were synthesized and tested for their antimitotic activity on a human breast cancer cell line against the prototype A10 and the antibreast cancer drug tamoxifen. Moreover, the DNA binding capacity of such compounds was evaluated against A10. (E)-3-(4-Nitrocinnamoylamino)-2,6-piperidinedione "3B" and (E)-3-(4-hydroxycinnamoylamino)-2,6-piperidinedione "3D" were several-fold more potent antiproliferative agents than A10 and tamoxifen. They also had significantly higher capacity to bind DNA than A10. Conversely, (E)-3-(cinnamoylamino)-2,6-piperidinedione "3A" and (E)-3-(4-methoxycinnamoylamino)-2,6-piperidinedione) "3C" had weaker biological profiles than the lead compound A10. Detailed synthetic, spectroscopic, and biological data are reported.

CONCEPT CODE:

Reproductive system - Physiology and biochemistry 16504

Cytology - Human 02508

Biochemistry studies - General 10060

Biochemistry studies - Nucleic acids, purines and

pyrimidines 10062

Pathology - Therapy 12512

Reproductive system - Pathology 16506

Pharmacology - General 22002

Pharmacology - Clinical pharmacology 22005

Neoplasms - Pathology, clinical aspects and systemic

effects 24004

Neoplasms - Therapeutic agents and therapy 24008

INDEX TERMS: Major Concepts

Pharmacology; Reproductive System (Reproduction); Tumor

Biology

INDEX TERMS: Diseases

breast cancer: neoplastic disease, reproductive system

disease

Breast Neoplasms (MeSH)

Chemicals & Biochemicals

INDEX TERMS:

(E) -3-(4-hydroxycinnamoylamino) -2,6-piperidinedione:

antimitotic-drug, antineoplastic-drug

, antimitotic activity, synthesis; (E)-3-(4methoxycinnamoylamino)-2,6-piperidinedione):

antimitotic-drug, antineoplastic-drug

, antimitotic activity, synthesis; (E)-3-(4-

nitrocinnamoylamino) -2,6-piperidinedione: antimitotic-

drug, antineoplastic-drug, antimitotic

activity, synthesis; (E)-3-(cinnamoylamino)-2,6-

piperidinedione: antimitotic-drug,

antineoplastic-drug, antimitotic activity,

synthesis; 3-phenylacetylamino-2,6-piperidinedione

[A-10]: antineoplastic-drug, analogs,

antineoplaston; DNA; piperidinedione analogs:

antineoplastic activity; tamoxifen: antineoplastic-

drug

INDEX TERMS:

Methods & Equipment

spectroscopic analysis: analytical method

INDEX TERMS:

Miscellaneous Descriptors

binding affinity

ORGANISM:

Classifier

Hominidae 86215

Super Taxa

Primates; Mammalia; Vertebrata; Chordata; Animalia

Organism Name

MCF-7 cell line: human breast cancer cells

Taxa Notes

Animals, Chordates, Humans, Mammals, Primates,

Vertebrates

REGISTRY NUMBER: 77658-84-5 (3-phenylacetylamino-2,6-

> piperidinedione) 77658-84-5 (A-10) 10540-29-1 (tamoxifen)

L131 ANSWER 34 OF 44 BIOSIS COPYRIGHT (c) 2005 The Thomson Corporation. on

STN

ACCESSION NUMBER: 1997:356904 BIOSIS DOCUMENT NUMBER: PREV199799663307

TITLE: Enantiomeric separation of some piperidine-2,6-dione drugs on tolylcellulose by liquid chromatography.

AUTHOR(S): Van Overbeke, An; Aboul-Enein, Hassan Y. [Reprint author];

Baeyens, Willy; Van Der Weken, Guido; Dewaele, Chris

Biological and Med. Res., MBC-03, King Faisal Specialist CORPORATE SOURCE:

Hosp. and Res. Centre, P.O. Box 33544, Riyadh 11211, Saudi

Arabia

SOURCE: Analytica Chimica Acta, (1997) Vol. 346, No. 2, pp.

183-189.

CODEN: ACACAM. ISSN: 0003-2670.

DOCUMENT TYPE:

Article LANGUAGE: English

Entered STN: 25 Aug 1997 **ENTRY DATE:**

Last Updated on STN: 27 Oct 1997

ABSTRACT: The direct liquid chromatographic resolution of a group of drugs, all having a piperidine-2,6-dione structure in common was investigated using a cellulose-based chiral stationary phase. The experimental tris(4methylbenzoate) cellulose column (BioRad RSL, Belgium) has the polymeric layer covalently bonded onto an aminopropylated silica support. Reversed phase as well as normal phase conditions were applied without deterioration of the Aminoglutethimide and glutethimide were easily resolved using methanol-water mixtures as mobile phase. Acetylaminoglutethimide, the major metabolite of aminoglutethimide, was separated from the parent drug but its enantiomers were not completely resolved. Thalidomide enantiomers were at best resolved under normal phase conditions with n-hexane as the major constituent of the mobile phase. No chiral interaction on this column was noticed for 3-phenylacetylaminopiperidine-2,6-dione.

CONCEPT CODE:

Biochemistry methods - General Biochemistry studies - General Biophysics - Methods and techniques

Pharmacology - General 22002

INDEX TERMS:

Major Concepts

Biochemistry and Molecular Biophysics; Methods and

Techniques; Pharmacology

INDEX TERMS:

Chemicals & Biochemicals

PIPERIDINE-2,6-DIONE; GLUTETHIMIDE; AMINOGLUTETHIMIDE; 3-PHENYLACETYLAMINOPIPERIDINE-2,6-DIONE; THALIDOMIDE

INDEX TERMS:

Miscellaneous Descriptors

AMINOGLUTETHIMIDE; CYCLOHEXYLAMINOGLUTETHIMIDE; ENANTIOMERIC SEPARATION; GLUTETHIMIDE; METHODOLOGY;

N-ACETYLAMINOGLUTETHIMIDE; NORMAL PHASE;

PHARMACEUTICALS; PIPERIDINE-2,6-DIONE DRUGS; REVERSED PHASE; SEPARATION METHOD; THALIDOMIDE; TOLYLCELLULOSE

COLUMN LIQUID CHROMATOGRAPHY; 3-PHENYLACETYLAMINOPIPERIDINE-2,6-DIONE

REGISTRY NUMBER:

1121-89-7 (PIPERIDINE-2,6-DIONE)

77-21-4 (GLUTETHIMIDE)

125-84-8 (AMINOGLUTETHIMIDE)

77658-84-5 (3-PHENYLACETYLAMINOPIPERIDINE-2,6-

Jones 09/955010

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DIONE)

50-35-1 (THALIDOMIDE)

L131 ANSWER 35 OF 44 BIOSIS COPYRIGHT (c) 2005 The Thomson Corporation. or

STN

ACCESSION NUMBER: 1993:390859 BIOSIS DOCUMENT NUMBER: PREV199396066159

TITLE: Further studies on the specificity of interaction of

3-phenylacetylamino-2,6-piperidinedione with DNA.

AUTHOR(S): Lehner, A. F. [Reprint author]; Burzynski, S. R.; Hendry,

L. B.

CORPORATE SOURCE: Augusta Lab. Inc., PO Box 3293, Augusta, GA 30904, USA

SOURCE: International Journal of Experimental and Clinical

Chemotherapy, (1992) Vol. 5, No. 2, pp. 63-71.

CODEN: IJECED. ISSN: 0933-0453.

DOCUMENT TYPE: Article LANGUAGE: English

ENTRY DATE: Entered STN: 23 Aug 1993

Last Updated on STN: 28 Sep 1993

ABSTRACT: Spectroscopic studies were carried out on the interactions of DNA with Antineoplaston A10 (3-phenylacetylamino-2,6-piperidinedione), a dipeptide with specific chemical modifications and with a described antineoplastic activity. DNA thermal denaturation studies indicated that A10 can interact with defined polydeoxynucleotides, as seen most notably in the concentration-dependent stabilization of poly(dAdG)*poly(dCdT). These findings agree with DNA modeling studies which have demonstrated that A10 is capable of stereospecific insertion between certain base pair sequences in DNA, in particular 5'-dTdT-3'*5'-dAdA-3',5'-dTdC-3'*5'-dGdA-3', and 5'-dCdT-3'*5'-dAdG-3'. Note that 2 of these two-base sequences occur repeatedly in poly(dAdG) *poly(dCdT). The effects of Alo on DNA were compared with those of the Alo hydrolysis product phenylacetylisoglutamine (PAisoG). The A10-DNA interaction was much stronger than that between PAisoG and DNA by the Tm criteria, and this suggests that the piperidinedione ring of A10 plays an essential role in stabilizing DNA during thermal denaturation. The latter concept was supported by observation of even weaker interactions of the DNA with phenylacetic acid, which is the residue after complete removal of the amino-2,6-piperidinedione moiety from A10. In light of these physicochemical studies and their general agreement with DNA modeling, a scheme is proposed whereby A10 may exert some of its antineoplastic properties by prevention or alleviation of certain mutagenic lesions at DNA sequences containing adjacent pyrimidines.

CONCEPT CODE: Biochemistry studies - Nucleic acids, purines and

pyrimidines 10062

Biochemistry studies - Proteins, peptides and amino acids

10064

Biophysics - Molecular properties and macromolecules

10506

Pharmacology - General 22002

Neoplasms - Therapeutic agents and therapy 24008

INDEX TERMS: Major Concepts

Biochemistry and Molecular Biophysics; Pharmacology;

Tumor Biology

INDEX TERMS: Chemicals & Biochemicals

3-PHENYLACETYLAMINO-2,6-PIPERIDINEDIONE; ANTINEOPLASTON

A10

INDEX TERMS: Miscellaneous Descriptors

ANTINEOPLASTIC-DRUG; TUMOR CELL DEVELOPMENT

REGISTRY NUMBER: 77658-84-5 (3-PHENYLACETYLAMINO-2,6-

PIPERIDINEDIONE)

91531-30-5 (ANTINEOPLASTON A10)

L131 ANSWER 36 OF 44 BIOSIS COPYRIGHT (c) 2005 The Thomson Corporation. on STN

ACCESSION NUMBER: 1991:480777 BIOSIS

DOCUMENT NUMBER: PREV199192114537; BA92:114537

3 PHENYLACETYLAMINO-2 6-PIPERIDINEDIONE INHIBITION OF RAT TITLE:

NB2 LYMPHOMA CELL MITOGENESIS.

AUTHOR(S): WOOD J C [Reprint author]; COPLAND J A; MULDOON T G; HENDRY

DEP PHYSIOL AND ENDOCRINOL, CLW 334, MEDICAL COLL GA, CORPORATE SOURCE:

AUGUSTA, GA 30912, USA

Proceedings of the Society for Experimental Biology and Medicine, (1991) Vol. 197, No. 4, pp. 404-408. SOURCE:

CODEN: PSEBAA. ISSN: 0037-9727.

DOCUMENT TYPE:

Article

FILE SEGMENT: LANGUAGE:

ENGLISH

BA

ENTRY DATE:

Entered STN: 26 Oct 1991

Last Updated on STN: 8 Jan 1992

ABSTRACT: 3-Phenylacetylamino-2,6-piperidinedione (A10), an amino acid analog, has been reported to possess antineoplastic activity against certain neoplastic tissues. The antimitogenic properties of A10 were studied by determining its effect on prolactin (PRL) - and interleukin 2 (IL-2)-stimulated mitogenic responses in the rat Nb2 lymphoma cell line. The addition of A10 (1-12 mM) to PRL (0.4 ng/ml)-stimulated cells inhibited growth in a dose-dependent manner. DNA synthesis patterns studies by thymidine incorporation demonstrated that A10 was significantly inhibitory (25% at 20 hr; 50% at 40 hr, P < 0.01). IL-2 stimulation of mitogenesis was also sensitive to A10 inhibition. inhibition of PRL stimulated mitogenesis was reversible when A10 was removed after 24 hr of culture and A10 showed no toxicity in a chromium release assay. These data suggest that A10 effects may be cytostatic, rather than cytotoxic.

CONCEPT CODE:

Cytology - Animal 02506

Biochemistry studies - Nucleic acids, purines and

pyrimidines 10062

Biochemistry studies - Proteins, peptides and amino acids

Biochemistry studies - Carbohydrates 10068

Pathology - Therapy 12512

Metabolism - Nucleic acids, purines and pyrimidines 13014

Blood - Blood, lymphatic and reticuloendothelial

pathologies 15006

Blood - Lymphatic tissue and reticuloendothelial system

15008

Endocrine - General 17002 Endocrine - Pituitary 17014

Pharmacology - Clinical pharmacology 22005 Pharmacology - Blood and hematopoietic agents 22008

Neoplasms - Neoplastic cell lines

Neoplasms - Therapeutic agents and therapy

Neoplasms - Blood and reticuloendothelial neoplasms 24010

INDEX TERMS: Major Concepts

Blood and Lymphatics (Transport and Circulation);

Endocrine System (Chemical Coordination and

Homeostasis); Metabolism; Pharmacology; Tumor Biology

Miscellaneous Descriptors INDEX TERMS:

ANTINEOPLASTIC-DRUG PROLACTIN INTERLEUKIN 2

DNA SYNTHESIS

ORGANISM:

Classifier

Muridae 86375

Super Taxa

Rodentia; Mammalia; Vertebrata; Chordata; Animalia

Taxa Notes

Animals, Chordates, Mammals, Nonhuman Vertebrates,

Nonhuman Mammals, Rodents, Vertebrates

77658-84-5 (3-PHENYLACETYLAMINO-2,6-REGISTRY NUMBER:

> PIPERIDINEDIONE) 9002-62-4 (PROLACTIN)

L131 ANSWER 37 OF 44 BIOSIS COPYRIGHT (c) 2005 The Thomson Corporation. on

STN

ACCESSION NUMBER: 1991:354729 BIOSIS

DOCUMENT NUMBER: PREV199141039244; BR41:39244

IN-VITRO AND IN-VIVO INHIBITION OF ESTROGEN INDUCED TUMOR TITLE:

GROWTH BY A NOVEL ANTITUMOR COMPOUND.

COPLAND J A [Reprint author]; WOOD J C; HENDRY L B; AUTHOR (S):

PANTAZIS C G; CHU C K; MAHESH V B

CORPORATE SOURCE: DEP PHYSIOL AND ENDOCRINOL, MED GA, AUGUSTA, GA 30912, USA

SOURCE: Proceedings of the American Association for Cancer Research

Annual Meeting, (1991) Vol. 32, pp. 131.

Meeting Info.: 82ND ANNUAL MEETING OF THE AMERICAN

ASSOCIATION FOR CANCER RESEARCH, HOUSTON, TEXAS, USA, MAY

15-18, 1991. PROC AM ASSOC CANCER RES ANNU MEET.

ISSN: 0197-016X.

DOCUMENT TYPE:

Conference; (Meeting)

FILE SEGMENT:

BR

ENGLISH LANGUAGE:

ENTRY DATE: Entered STN: 1 Aug 1991

Last Updated on STN: 11 Sep 1991

CONCEPT CODE: General biology - Symposia, transactions and proceedings

00520

Cytology - Animal 02506 Cytology - Human 02508

Biochemistry studies - General 10060

Biochemistry studies - Sterols and steroids 10067

Pathology - Therapy 12512 Endocrine - Gonads and placenta 17006 Pharmacology - General Pharmacology - Endocrine system 22016

Neoplasms - Therapeutic agents and therapy 24008

INDEX TERMS: Major Concepts

Cell Biology; Oncology (Human Medicine, Medical

Sciences); Pharmacology

INDEX TERMS: Miscellaneous Descriptors

ABSTRACT HUMAN MCF-7 CELLS MOUSE 3 PHENYLACETYLAMINO-2

6-PIPERIDINEDIONE P HYDROXY-3-PHENYLACETYLAMINO-2

6-PIPERIDINEDIONE ANTINEOPLASTIC-DRUG

ANTIESTROGEN AGENTS

Classifier ORGANISM:

> Hominidae 86215

Super Taxa

Primates; Mammalia; Vertebrata; Chordata; Animalia

Taxa Notes

Animals, Chordates, Humans, Mammals, Primates,

Vertebrates

Classifier ORGANISM:

> Muridae 86375

Super Taxa

Rodentia; Mammalia; Vertebrata; Chordata; Animalia

Taxa Notes

Animals, Chordates, Mammals, Nonhuman Vertebrates,

Nonhuman Mammals, Rodents, Vertebrates

REGISTRY NUMBER: 77658-84-5 (3-PHENYLACETYLAMINO-2

6-PIPERIDINEDIONE)

L131 ANSWER 38 OF 44 BIOSIS COPYRIGHT (c) 2005 The Thomson Corporation. on STN

ACCESSION NUMBER: 1990:348286 BIOSIS

DOCUMENT NUMBER: PREV199039043547; BR39:43547

TITLE: PARA-HYDROXYLATION OF 3 PHENYLACETYLAMINO-2

6-PIPERIDINEDIONE INCREASES THE INHIBITION OF PROLACTIN

STIMULATED NB-2 CELL MITOGENESIS.

WOOD J C [Reprint author]; HUANG H Q; CHU C K; HENDRY L B AUTHOR (S):

DEP PHYSIOL AND ENDOCRINOL, MED COLL GA, AUGUSTA, GA 30912, CORPORATE SOURCE:

SOURCE: Proceedings of the American Association for Cancer Research

Annual Meeting, (1990) Vol. 31, pp. 410.

Meeting Info.: 81ST ANNUAL MEETING OF THE AMERICAN

ASSOCIATION FOR CANCER RESEARCH, WASHINGTON, D.C., USA, MAY

23-26, 1990. PROC AM ASSOC CANCER RES ANNU MEET.

ISSN: 0197-016X.

DOCUMENT TYPE: Conference; (Meeting)

FILE SEGMENT:

LANGUAGE: **ENGLISH**

Entered STN: 26 Jul 1990 ENTRY DATE:

Last Updated on STN: 30 Aug 1990

CONCEPT CODE: General biology - Symposia, transactions and proceedings

00520

Cytology - Animal 02506 Genetics - Animal 03506

Biochemistry studies - General

Biochemistry studies - Nucleic acids, purines and

pyrimidines 10062

Pathology - Therapy 12512

Metabolism - General metabolism and metabolic pathways

Pharmacology - Drug metabolism and metabolic stimulators

22003

Neoplasms - Neoplastic cell lines

Neoplasms - Therapeutic agents and therapy 24008 In vitro cellular and subcellular studies

INDEX TERMS: Major Concepts

Genetics; Metabolism; Pharmacology; Tumor Biology

INDEX TERMS: Miscellaneous Descriptors

ABSTRACT ANTINEOPLASTIC-DRUG PHARMACOKINETICS

DNA BINDING

ORGANISM:

Classifier

Muridae 86375

Super Taxa

Rodentia; Mammalia; Vertebrata; Chordata; Animalia

Taxa Notes

Animals, Chordates, Mammals, Nonhuman Vertebrates,

Nonhuman Mammals, Rodents, Vertebrates

REGISTRY NUMBER:

77658-84-5 (3-PHENYLACETYLAMINO-2,6-

PIPERIDINEDIONE) 9002-62-4 (PROLACTIN)

L131 ANSWER 39 OF 44 BIOSIS COPYRIGHT (c) 2005 The Thomson Corporation. on

STN

ACCESSION NUMBER: 1990:467114 BIOSIS

DOCUMENT NUMBER: PREV199039102475; BR39:102475

TITLE: STRUCTURE-ACTIVITY RELATIONSHIPS MOLECULAR MODELING AND

ANTITUMOR ACTIVITY OF 2 6 PIPERIDINEDIONES.

AUTHOR (S): HENDRY L B [Reprint author]; CHU C K; HUANG H; WOOD J C;

COPLAND J A; MAHESH V B

CORPORATE SOURCE: DEP PHYSIOL ENDOCRINOL, MEDICAL COLLEGE GEORGIA,

STEREOCHEMICAL GENETICS INC, PO BOX 11649, AUGUSTA, GA

30912, USA

SOURCE: Abstracts of Papers American Chemical Society, (1990) Vol.

200, No. 1-2, pp. MEDI 148.

Meeting Info.: 200TH AMERICAN CHEMICAL SOCIETY NATIONAL MEETING, WASHINGTON, D.C., USA, AUGUST 26-31, 1990. ABSTR

PAP AM CHEM SOC.

CODEN: ACSRAL. ISSN: 0065-7727.

DOCUMENT TYPE:

Conference; (Meeting)

FILE SEGMENT:

BR

LANGUAGE:

ENGLISH

ENTRY DATE:

Entered STN: 13 Oct 1990

Last Updated on STN: 4 Jan 1991

CONCEPT CODE:

General biology - Symposia, transactions and proceedings

00520

Biochemistry studies - General 10060

Biochemistry studies - Nucleic acids, purines and

pyrimidines 10062

Biophysics - Molecular properties and macromolecules

10506

Pharmacology - Clinical pharmacology 22005

Neoplasms - Biochemistry 24006

Neoplasms - Therapeutic agents and therapy 24008

INDEX TERMS:

Major Concepts

Biochemistry and Molecular Biophysics; Oncology (Human

Medicine, Medical Sciences); Pharmacology

INDEX TERMS:

Miscellaneous Descriptors

ABSTRACT HUMAN RAT MOUSE 3 PHENYLACETYLAMINO-2 6-PIPERIDINEDIONE ANTINEOPLASTIC-DRUG DNA

COMPLEX

ORGANISM:

Classifier

Hominidae 86215

Super Taxa

Primates; Mammalia; Vertebrata; Chordata; Animalia

Taxa Notes

Animals, Chordates, Humans, Mammals, Primates,

Vertebrates

ORGANISM:

Classifier

Muridae 86375

Super Taxa

Rodentia; Mammalia; Vertebrata; Chordata; Animalia

Taxa Notes

Animals, Chordates, Mammals, Nonhuman Vertebrates,

Nonhuman Mammals, Rodents, Vertebrates

REGISTRY NUMBER:

1121-89-7D (2,6-PIPERIDINEDIONES)

77658-84-5 (3-PHENYLACETYLAMINO-2

6-PIPERIDINEDIONE)

L131 ANSWER 40 OF 44 BIOSIS COPYRIGHT (c) 2005 The Thomson Corporation. on

STN

ACCESSION NUMBER:

1989:20538 BIOSIS

DOCUMENT NUMBER:

PREV198936008215; BR36:8215

TITLE: ACTIONS OF AN ENDOGENOUS ANTITUMORIGENIC AGENT ON MAMMARY

TUMOR DEVELOPMENT AND MODELING ANALYSIS OF ITS CAPACITY FOR

INTERACTING WITH DNA.

AUTHOR(S):

HENDRY L B [Reprint author]; MULDOON T G

CORPORATE SOURCE:

DEP MED, MED COLL GEORGIA, AUGUSTA, GA 30912, USA

SOURCE:

Journal of Steroid Biochemistry, (1988) Vol. 30, No. 1-6,

pp. 325-328.

Meeting Info.: MEETING ON RECENT ADVANCES IN STEROID

BIOCHEMISTRY HELD AT THE EIGHTH INTERNATIONAL SYMPOSIUM OF THE JOURNAL OF STEROID BIOCHEMISTRY, PARIS, FRANCE, MAY

24-27, 1987. J STEROID BIOCHEM. CODEN: JSTBBK. ISSN: 0022-4731.

DOCUMENT TYPE:

Conference; (Meeting)

09/955010 Page 38 Jones

FILE SEGMENT:

BR

LANGUAGE:

ENGLISH

Entered STN: 13 Dec 1988

ENTRY DATE:

Last Updated on STN: 13 Dec 1988

CONCEPT CODE:

General biology - Symposia, transactions and proceedings

00520

Genetics - Animal

03506 Biochemistry studies - Nucleic acids, purines and

pyrimidines 10062

Reproductive system - Pathology 16506 Endocrine - Gonads and placenta 17006 Pharmacology - Endocrine system 22016

Neoplasms - Biochemistry 24006

Neoplasms - Therapeutic agents and therapy 24008

INDEX TERMS:

Major Concepts

Endocrine System (Chemical Coordination and

Homeostasis); Genetics; Pharmacology; Reproductive

System (Reproduction); Tumor Biology

INDEX TERMS:

Miscellaneous Descriptors RAT MOUSE ANTINEOPLASTIN A-10 3 PHENYLACETYLAMINO-2 6-PIPERIDINEDIONE ANTINEOPLASTIC-DRUG ANDROGEN

INHIBITION ANTI-ESTROGEN SUPPRESSION

ORGANISM:

Classifier

Muridae 86375

Super Taxa

Rodentia; Mammalia; Vertebrata; Chordata; Animalia

Taxa Notes

Animals, Chordates, Mammals, Nonhuman Vertebrates,

Nonhuman Mammals, Rodents, Vertebrates

REGISTRY NUMBER:

77658-84-5 (3-PHENYLACETYLAMINO-2

6-PIPERIDINEDIONE)

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STN

ACCESSION NUMBER:

1986:288757 BIOSIS

DOCUMENT NUMBER:

PREV198631023335; BR31:23335

TITLE:

TOPICAL USE OF 3 PHENYLACETYLAMINO-2 6-PIPERIDINEDIONE FOR

TREATMENT OF SKIN WRINKLES AND HYPERPIGMENTATION US

PATENT-4593038. JUNE 3 1986.

AUTHOR(S): CORPORATE SOURCE: BURZYNSKI S R [Inventor, Reprint author] 5 CONCORD CIR, HOUSTON, TEX 77024, USA

PATENT INFORMATION: US 4593038 June 03, 1986

SOURCE:

Official Gazette of the United States Patent and Trademark

Office Patents, (1986) Vol. 1067, No. 1, pp. 318.

CODEN: OGUPE7. ISSN: 0098-1133.

DOCUMENT TYPE:

Patent BR

FILE SEGMENT: LANGUAGE:

ENGLISH

ENTRY DATE:

Entered STN: 12 Jul 1986

Last Updated on STN: 12 Jul 1986

NAT. PATENT. CLASSIF.:514328000

CONCEPT CODE:

Biochemistry studies - General 10060

Pathology - Therapy 12512

Integumentary system - General and methods

Pharmacology - Clinical pharmacology

Pharmacology - Integumentary system, dental and oral

biology 22020

Routes of immunization, infection and therapy 22100

INDEX TERMS: Major Concepts

Biochemistry and Molecular Biophysics; Integumentary

System (Chemical Coordination and Homeostasis);

Pharmacology

INDEX TERMS:

Miscellaneous Descriptors

USCL-514-328 DERMATOLOGICAL-DRUG

REGISTRY NUMBER:

77658-84-5 (3-PHENYLACETYLAMINO-2,6-

PIPERIDINEDIONE)

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STN

ACCESSION NUMBER:

1987:79848 BIOSIS

DOCUMENT NUMBER:

PREV198732040041; BR32:40041

TITLE:

INVESTIGATION OF THE MODE OF ACTION OF ANTINEOPLASTON A10

EVIDENCE FOR BINDING TO DNA.

AUTHOR(S):

HENDRY L B [Reprint author]; LEHNER A F; MULDOON T G;

COPLAND J A; MAHESH V B; MILL T M; BURZYNSKI S R

CORPORATE SOURCE:

MED COLL GA, AUGUSTA, GA 30912, USA

SOURCE:

(1986) pp. 607. UICC (UNION INTERNATIONALE CONTRE LE CANCER, INTERNATIONAL UNION AGAINST CANCER). 14TH INTERNATIONAL CANCER CONGRESS, BUDAPEST, HUNGARY, AUG. 21-27, 1986. ABSTRACTS, LECTURES, SYMPOSIA AND FREE COMMUNICATIONS, VOLS. 1, 2, 3, LATE ABSTRACTS, AND REGISTER. XVI+479P. (VOL. 1); XVI+298P. (VOL. 2);

XVI+531P.(VOL. 3); 15P.(LATE ABSTRACTS); 40P.(REGISTER) S.

KARGER AG: BASEL, SWITZERLAND; NEW YORK, N.Y., USA;

AKADEMIAI KIADO: BUDAPEST, HUNGARY. PAPER.

ISBN: 3-8055-4434-0(KARGER), 963-05-4422-9(VOL. 1),

963-05-4423-7 (VOL. 2), 963-05-4424-5 (VOL. 3),

963-05-4439-3 (LATE ABSTRACTS), 963-05-4425-3 (REGISTER),

963-05-4421-0 (GENERAL) .

DOCUMENT TYPE:

Book

Conference; (Meeting)

FILE SEGMENT:

BR ENGLISH

LANGUAGE: ENTRY DATE:

Entered STN: 31 Jan 1987

CONCEPT CODE:

Last Updated on STN: 31 Jan 1987 General biology - Symposia, transactions and proceedings

00520

Biochemistry studies - General 10060

Biochemistry studies - Nucleic acids, purines and

pyrimidines 10062

Biochemistry studies - Sterols and steroids 10067

Metabolism - Sterols and steroids 13008 Reproductive system - Pathology 16506

Pharmacology - General 22002

Pharmacology - Reproductive system and implantation studies

22028

Neoplasms - Biochemistry 24006

Neoplasms - Therapeutic agents and therapy In vitro cellular and subcellular studies 32600

INDEX TERMS:

Major Concepts

Metabolism; Pharmacology; Reproductive System

(Reproduction); Tumor Biology

INDEX TERMS:

Miscellaneous Descriptors

ABSTRACT RAT 3 PHENYLACETYLAMINO-2 6-PIPERIDINEDIONE

ANTINEOPLASTIC-DRUG ESTROGEN METABOLISM BREAST

CARCINOMA

ORGANISM:

Muridae 86375

Super Taxa

Rodentia; Mammalia; Vertebrata; Chordata; Animalia

Taxa Notes

Classifier

Animals, Chordates, Mammals, Nonhuman Vertebrates,

Nonhuman Mammals, Rodents, Vertebrates

REGISTRY NUMBER:

91531-30-5 (ANTINEOPLASTON A10)

77658-84-5 (3-PHENYLACETYLAMINO-2 6-PIPERIDINEDIONE)

L131 ANSWER 43 OF 44 BIOSIS COPYRIGHT (c) 2005 The Thomson Corporation. on

STN

ACCESSION NUMBER: 1986:399248 BIOSIS

DOCUMENT NUMBER: PREV198682084728; BA82:84728

TITLE: 3 PHENYLACETYLAMINO-2 6-PIPERIDINEDIONE A

NATURALLY-OCCURRING PEPTIDE ANALOGUE WITH APPARENT

ANTINEOPLASTIC ACTIVITY MAY BIND TO DNA.

AUTHOR(S): LEHNER A F [Reprint author]; BURZYNSKI S R; HENDRY L B

CORPORATE SOURCE: DEP MED, MED COLLEGE GEORGIA, AUGUSTA, GEORGIA 30912, USA SOURCE: Drugs under Experimental and Clinical Research, (1986) Vol.

12, No. SUPPL. 1, pp. 57-72. CODEN: DECRDP. ISSN: 0378-6501.

DOCUMENT TYPE: Article

FILE SEGMENT: BA
LANGUAGE: ENGLISH

ENTRY DATE: Entered STN: 4 Oct 1986

Last Updated on STN: 4 Oct 1986

ABSTRACT: Antineoplaston A10 (3-phenylacetylamino-2,6-piperidinedione), a peptide analgoue originally isolated from human urine and serum, appears to have antineoplastic activity. In view of the close resemblance of the structure of AlO to that of DNA intercalative anticancer drugs, spectroscopic studies were performed to determine whether its mode of action could similarly involve binding to DNA. DNA thermal denaturation studies demonstrated that A10 was capable of interacting with DNA in a specific manner; of the synthetic polynucleotides employed in this study, AlO had the greatest effect on poly(dA-Dg) · poly(dC-dT), suggesting some sequence preference. However, ultraviolet and fluorescence spectroscopic studies demonstrated that interactions of A10 with DNA were weak in comparison to those of classical intercalating agents. Mass spectroscopic studies suggested that A10 did not react covalently with DNA. The weak yet apparently specific interaction of A10 with DNA indicates that the mode of action of A10 may involve binding to chromatin, facilitated by nuclear protein receptors analogous to steroid and thyroid hormones.

CONCEPT CODE:

Genetics - Animal 03506

Clinical biochemistry - General methods and applications

10006

Biochemistry methods - Nucleic acids, purines and

pyrimidines 10052

Biochemistry studies - Nucleic acids, purines and

pyrimidines 10062

Biochemistry studies - Proteins, peptides and amino acids

10064

Biochemistry studies - Sterols and steroids 10067

Biophysics - Methods and techniques 105 Biophysics - Membrane phenomena 10508 Blood - Blood and lymph studies 15002

Blood - Other body fluids 15010

Urinary system - Physiology and biochemistry 15504

Endocrine - Adrenals 17004 Endocrine - Thyroid 17018

Pharmacology - Drug metabolism and metabolic stimulators

22003

Neoplasms - Therapeutic agents and therapy 24008

INDEX TERMS: Major Concepts

Biochemistry and Molecular Biophysics; Genetics;

Pharmacology; Tumor Biology

INDEX TERMS: Miscellaneous Descriptors

ANTINEOPLASTON A-10 ANTINEOPLASTIC-DRUG

PHARMACODYNAMICS

REGISTRY NUMBER: 77658-84-5 (3-PHENYLACETYLAMINO-2,6-

PIPERIDINEDIONE)

91531-30-5 (ANTINEOPLASTON A-10)

L131 ANSWER 44 OF 44 TOXCENTER COPYRIGHT 2005 ACS on STN

ACCESSION NUMBER:

1984:715 TOXCENTER Copyright 2005 ASHP

COPYRIGHT: DOCUMENT NUMBER:

22-01046

TITLE:

Animal toxicology studies on oral formulation of

antineoplaston A10

AUTHOR(S):

Burzynski, S. R.; Mohabbat, M. O.; Burzynski, B.

CORPORATE SOURCE:

Burzynski Res. Inst., 12707 Trinity Dr., Stafford, TX

77477

SOURCE:

Drugs Under Experimental and Clinical Research (Switzerland), (1984) Vol. 10, pp. 113-118. 8 Refs.

CODEN: DECRDR. ISSN: 0378-6501.

DOCUMENT TYPE: FILE SEGMENT:

Journal IPA

OTHER SOURCE:

IPA 84:2496

LANGUAGE:

English

ENTRY DATE:

Entered STN: 20011116

Last Updated on STN: 20011116

ABSTRACT:

The toxic effects of antineoplaston A10

(3-phenylacetylamino-2,6-piperidimedione) were evaluated in mice. No

toxic effects were associated with the daily chronic oral

administration of the drug.

D. L. Thompson

SECTION CODE:

5 Investigational Drugs

CLASSIFICATION CODE: 10:00 Antineoplastic agents

SUPPLEMENTARY TERMS: Miscellaneous Descriptors

Antineoplaston A10; toxicity; lack, mice

Antineoplastic agents; antineoplaston A10; lacks

toxicity, mice

Toxicity; antineoplaston AlO; lack, mice

REGISTRY NUMBER:

77658-84-5 (Antineoplaston A10)

CHEMICAL NAME:

Antineoplaston A10 (3-Phenylacetylamino-2,6-

piperidinedione)

=> fil capl; d que 140

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FILE COVERS 1907 - 7 Jan 2005 VOL 142 ISS 3 FILE LAST UPDATED: 6 Jan 2005 (20050106/ED)

This file contains CAS Registry Numbers for easy and accurate substance identification.

'OBI' IS DEFAULT SEARCH FIELD FOR 'CAPLUS' FILE

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            2 SEA FILE=REGISTRY ABB=ON
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^{18}
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             1 SEA FILE=REGISTRY ABB=ON CITRULLINE/CN
L10
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                                        VALINE/CN
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L18
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                                       (L8 OR L9 OR L10)
        109418 SEA FILE=CAPLUS ABB=ON (L11 OR L12 OR L13 OR L14 OR L15 OR
L19
               L16)
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L25
          2177 SEA FILE=CAPLUS ABB=ON
                                      L18(L) (PAC OR DMA OR THU OR PKT)/RL
L26
L27
          2610 SEA FILE=CAPLUS ABB=ON
                                      L19(L) (PAC OR DMA OR THU OR PKT)/RL
            73 SEA FILE=CAPLUS ABB=ON
                                      L25 AND L26 AND L27
L28
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L30
         110024 SEA FILE=CAPLUS ABB=ON ANTITUMOR AGENTS/CT
L31
L34
         281794 SEA FILE=CAPLUS ABB=ON
                                      TOXICITY/OBI OR CYTOTOXICITY/OBI
         77032 SEA FILE=CAPLUS ABB=ON NEOPLASM INHIBITORS/CT
L35
L39
         16534 SEA FILE=CAPLUS ABB=ON
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               2A) EFFECT#/OBI
.<u>L.4.0</u>
            AND L28
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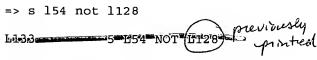
FILE USPATFULL ENTERED AT 12:30:22 ON 07 JAN 2005
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FILE COVERS 1971 TO PATENT PUBLICATION DATE: 6 Jan 2005 (20050106/PD)
FILE LAST UPDATED: 6 Jan 2005 (20050106/ED)
HIGHEST GRANTED PATENT NUMBER: US6839903
HIGHEST APPLICATION PUBLICATION NUMBER: US2005005336
CA INDEXING IS CURRENT THROUGH 6 Jan 2005 (20050106/UPCA)
ISSUE CLASS FIELDS (/INCL) CURRENT THROUGH: 6 Jan 2005 (20050106/PD)
REVISED CLASS FIELDS (/NCL) LAST RELOADED: Oct 2004
USPTO MANUAL OF CLASSIFICATIONS THESAURUS ISSUE DATE: Oct 2004
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>>> USPAT2 is now available. USPATFULL contains full text of the
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>>> applications. USPAT2 contains full text of the latest US
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>>> publications, starting in 2001, for the inventions covered in
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>>> published document but also a list of any subsequent
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>>> publications. The publication number, patent kind code, and
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>>> publication date for all the US publications for an invention
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>>> are displayed in the PI (Patent Information) field of USPATFULL
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>>> the earliest to the latest publication.
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This file contains CAS Registry Numbers for easy and accurate substance identification.

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             2 SEA FILE=REGISTRY ABB=ON ARGININE/CN
L8
             2 SEA FILE=REGISTRY ABB=ON ORNITHINE/CN
L9
            1 SEA FILE=REGISTRY ABB=ON CITRULLINE/CN
L10
            2 SEA FILE=REGISTRY ABB=ON ALANINE/CN
L11
            1 SEA FILE=REGISTRY ABB=ON GLYCINE/CN
L12
            2 SEA FILE=REGISTRY ABB=ON SERINE/CN
L13
            1 SEA FILE=REGISTRY ABB=ON
L14
                                        TAURINE/CN
            2 SEA FILE=REGISTRY ABB=ON
L15
                                        THREONINE/CN
            2 SEA FILE=REGISTRY ABB=ON
L16
                                        VALINE/CN
          994 SEA FILE=USPATFULL ABB=ON L7
L47
         2497 SEA FILE=USPATFULL ABB=ON (L8 OR L9 OR L10)
L48
L49
          6430 SEA FILE-USPATFULL ABB=ON (L11 OR L12 OR L13 OR L14 OR L15 OR
               L16)
           115 SEA FILE-USPATFULL ABB-ON L47 AND L48 AND L49
L51
L52
           538 SEA FILE=USPATFULL ABB=ON ((SIDE OR ADVERSE OR TOXIC)(2A)EFFEC
               T#)/IT
L53
          4703 SEA FILE-USPATFULL ABB-ON (TOXICITY OR CYTOTOXICITY)/IT
      6-SEA FILE USPATFULL ABBEON L51 AND (L52-OR L58)
L5.4
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=> fil biosis; d que 179; d que 180

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FILE COVERS 1969 TO DATE.

CAS REGISTRY NUMBERS AND CHEMICAL NAMES (CNs) PRESENT FROM JANUARY 1969 TO DATE.

RECORDS LAST ADDED: 5 January 2005 (20050105/ED)

FILE RELOADED: 19 October 2003.

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L7
                  1 SEA FILE=REGISTRY ABB=ON RIBOFLAVIN/CN
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L79 2=SEA=FILE=BIOSIS=ABB=ON—L74—AND—L76—AND—L78——
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                       OR L64)
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L134 2 L79 NOT (L129) mevicusty printed

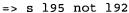
=> fil ipa; d que 195

CFILE TIPA ENTERED AT 12:30:25 ON 07 JAN 2005
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FILE COVERS 1970 TO 4 JAN 2005 (20050104/ED)

This file contains CAS Registry Numbers for easy and accurate substance identification.

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4£9.5≕		ESEA-	FILE=TPA-ABB=ON-(L82-OR-L83)=AND-(L84-OR-L85)-AND-(L86-OR.
		∠Lī8:7-	



L135 1-L95-NOT (L92) previously printed

=> fil toxcenter; d que l111

FILE 'TOXGENTER'-ENTERED AT 12:30:27 ON 07 JAN 2005 COPYRIGHT (C) 2005 ACS

FILE COVERS 1907 TO 4 Jan 2005 (20050104/ED)

This file contains CAS Registry Numbers for easy and accurate substance identification.

TOXCENTER has been enhanced with new files segments and search fields. See HELP CONTENT for more information.

TOXCENTER thesauri in the /CN, /CT, and /MN fields incorporate the MeSH 2005 vocabulary. See http://www.nlm.nih.gov/mesh/ and http://www.nlm.nih.gov/pubs/techbull/nd03/nd03_mesh.html for a description of changes.

L7 1 SEA FILE=REGISTRY ABB=ON RIBOFLAVIN/CN L8 2 SEA FILE=REGISTRY ABB=ON ARGININE/CN

L9	2	SEA FILE=REGISTRY ABB=ON ORNITHINE/CN
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		TAURINE OR THREONINE OR VALINE
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		T#)
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L107	2313689	SEA FILE=TOXCENTER ABB=ON TOXIC? OR CYTOTOXIC?
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L1-1-1-	10	SEA-FILE=TOXCENTER-ABB=ON-L110-AND-(AMINO-OR-COMPOSITION-OR-
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FILE 'WPIDS' ENTERED AT 12:30:28 ON 07 JAN 2005 COPYRIGHT (C) 2005 THE THOMSON CORPORATION

FILE LAST UPDATED: 3 JAN 2005 <20050103/UP>
MOST RECENT DERWENT UPDATE: 200501 <200501/DW>
DERWENT-WORLD-PATENTS-INDEX_SUBSCRIBER FILE, COVERS 1963 TO DATE

>>> FOR A COPY OF THE DERWENT WORLD PATENTS INDEX STN USER GUIDE, PLEASE VISIT:

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- >>> SMILES and ISOSMILES strings are no longer available as

Jones 09/955010

Page 47

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http://thomsonderwent.com/support/dwpiref/reftools/classification/code-revision/ FOR DETAILS. <<<</pre>

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               B2 OR B 2)
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FILE 'USPATFULL' ENTERED AT 12:31:22 ON 07 JAN 2005
CA INDEXING COPYRIGHT (C) 2005 AMERICAN CHEMICAL SOCIETY (ACS)

FILE 'IPA' ENTERED AT 12:31:22 ON 07 JAN 2005 COPYRIGHT (C) 2005 American Society of Hospital Pharmacists (ASHP)

FILE 'BIOSIS' ENTERED AT 12:31:22 ON 07 JAN 2005 Copyright (c) 2005 The Thomson Corporation.

FILE 'TOXCENTER' ENTERED AT 12:31:22 ON 07 JAN 2005 COPYRIGHT (C) 2005 ACS

L137—33—DUP_REM_L132_L133_L135_L134_L13.6_L123_(1_DUPLTCATE_REMOVED)

ANSWER '1' FROM FILE CAPLUS

ANSWERS '2-6' FROM FILE USPATFULL

ANSWER '7' FROM FILE IPA

ANSWERS '8-9' FROM FILE BIOSIS

ANSWERS '10-18' FROM FILE TOXCENTER

ANSWERS '19-33' FROM FILE WPIDS

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L137 ANSWER 1 OF 33 CAPLUS COPYRIGHT 2005 ACS on STN

ACCESSION NUMBER: 2000:608584 CAPLUS

DOCUMENT NUMBER: 133:187987

TITLE: Methods using pyrimidine-based nucleosides for

treatment of mitochondrial disorders

INVENTOR(S): Naviaux, Robert K.

PATENT ASSIGNEE(S): The Regents of the University of California, USA

SOURCE: PCT Int. Appl., 28 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

	PATENT NO.						KIND DATE				APPLICATION NO.						DATE				
	WO	2000	05004	43		A1 20000831			WO 2000-US4663					20000223							
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			IN,	IS,	JP,	KΕ,	KG,	ΚP,	KR,	KZ,	LC	, LK,	LR,	LS,	LT,	LU,	LV,	MA,			
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			SK,	SL,	ТJ,	TM,	TR,	TT,	TZ,	UA,	UG	, US,	UZ,	VN,	ΥU,	ZA,	ZW,	AM,			
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			ΙE,	SI,	LT,	LV,	FI,	RO													
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			•								WO :	2000-1	US46	63	1	₩ 2	0000	223			
											US :	2001-	8892	51		A1 2	0011	101			

OTHER SOURCE(S): MARPAT 133:187987

ED Entered STN: 01 Sep 2000

AB Methods are provided for the treatment of mitochondrial disorders. The methods include the administration of a pyrimidine-based nucleoside, e.g. triacetyluridine. Also provided are methods of reducing or eliminating symptoms associated with mitochondrial disorders. Mitochondrial disorders particularly appropriate for treatment include those attributable to a deficiency of one or more pyrimidines.

- IC ICM A61K031-70
- CC 1-12 (Pharmacology)
- IT Toxicity

(drug; pyrimidine-based nucleoside for treatment of mitochondrial disorder)

IT Antitumor agents

(leukemia, thrombocytopenia and leukemia syndrome; pyrimidine-based nucleoside for treatment of mitochondrial disorder)

IT Antitumor agents

(spleen lymphoma; pyrimidine-based nucleoside for treatment of mitochondrial disorder)

IT 51-35-4D, L-Hydroxyproline, pyrimidine nucleoside derivs. 52-90-4D,
L-Cysteine, pyrimidine nucleoside derivs., biological studies
56-40-6D, Glycine, pyrimidine nucleoside derivs., biological
studies 56-41-7D, L-Alanine, pyrimidine nucleoside derivs.,
biological studies 56-45-1D, L-Serine, pyrimidine nucleoside
derivs., biological studies 56-84-8D, L-Aspartic acid, pyrimidine

nucleoside derivs., biological studies 56-86-0D, L-Glutamic acid, pyrimidine nucleoside derivs., biological studies 56-87-1D, L-Lysine, pyrimidine nucleoside derivs., biological studies 56-89-3D, L-Cystine, pyrimidine nucleoside derivs., biological studies 58-85-5, Biotin 59-43-8, Vitamin B1, biological 59-30-3, Folic acid, biological studies 59-67-6, Niacin, biological studies 60-18-4D, L-Tyrosine, pyrimidine nucleoside derivs., biological studies 61-90-5D, L-Leucine, pyrimidine nucleoside derivs., biological studies 68-19-9, Vitamin B12 71-00-1D, 70-26-8D, L-Ornithine, pyrimidine nucleoside derivs. L-Histidine, pyrimidine nucleoside derivs., biological studies 72-18-4D, L-Valine, pyrimidine nucleoside derivs., biological studies **72-19-5D**, L-Threonine, pyrimidine nucleoside derivs., biological studies 73-32-5D, L-Isoleucine, pyrimidine nucleoside derivs., biological studies 74-79-3D, L-Arginine, pyrimidine nucleoside derivs., biological studies 79-83-4, Pantothenic acid 83-88-5, Vitamin B2, biological studies 147-85-3D, L-Proline, pyrimidine nucleoside derivs., biological studies 541-15-1D, L-Carnitine, pyrimidine nucleoside derivs. 4105-38-8 8059-24-3, Vitamin B6 52009-14-0, Calcium pyruvate RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(pyrimidine-based nucleoside for treatment of mitochondrial disorder)

REFERENCE COUNT: 2 THERE ARE 2 CITED REFERENCES AVAILABLE FOR THIS

RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L137 ANSWER 2 OF 33 USPATFULL on STN

ACCESSION NUMBER: 2004:286752 USPATFULL

TITLE: Methods of treatment of mitochondrial disorders
INVENTOR(S): Naviaux, Robert K., San Diego, CA, UNITED STATES

PATENT ASSIGNEE(S): THE REGENTS OF THE UNIVERSITY OF CALIFORNIA (U.S.

corporation)

RELATED APPLN. INFO.: Continuation of Ser. No. US 2001-889251, filed on 1 Nov

2001, PENDING A 371 of International Ser. No. WO

2000-US4663, filed on 23 Feb 2000, PENDING

NUMBER DATE

PRIORITY INFORMATION: US 1999-121588P 19990223 (60)

DOCUMENT TYPE: Utility
FILE SEGMENT: APPLICATION

LEGAL REPRESENTATIVE: Lisa A. Haile, J.D., Ph.D., GRAY CARY WARE &

FREIDENRICH LLP, Suite 1100, 4365 Executive Drive, San

Diego, CA, 92121-2133

NUMBER OF CLAIMS: 28
EXEMPLARY CLAIM: CLM-01-27
LINE COUNT: 716

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

AB In accordance with the present invention, there are provided methods for the treatment of mitochondrial disorders. Invention methods include the administration of a pyrimidine-based nucleoside such as triacetyluridine, or the like. Also provided are methods of reducing or eliminating symptoms associated with mitochondrial disorders.

Mitochondrial disorders particularly appropriate for treatment include

those attributable to a deficiency of one or more pyrimidines.

56-40-6D, Glycine, pyrimidine nucleoside derivs., biological studies 56-41-7D, L-Alanine, pyrimidine nucleoside derivs., biological studies 56-45-1D, L-Serine, pyrimidine nucleoside derivs., biological studies 70-26-8D, L-Ornithine, pyrimidine nucleoside derivs., biological studies 70-26-8D, L-Threonine, pyrimidine nucleoside derivs., biological studies 72-19-5D, L-Threonine, pyrimidine nucleoside derivs., biological studies 74-79-3D, L-Arginine, pyrimidine nucleoside derivs., biological studies 83-88-5, Vitamin B2, biological studies

(pyrimidine-based nucleoside for treatment of mitochondrial disorder)

L137 ANSWER 3 OF 33 USPATFULL on STN

ACCESSION NUMBER: 2004:50763 USPATFULL

TITLE: System for exsanguinous metabolic support of an organ

or tissue

INVENTOR(S): Brasile, Lauren, Albany, NY, UNITED STATES

PATENT ASSIGNEE(S): Breonics, Inc., Otisville, NY (U.S. corporation)

NUMBER KIND DATE

PATENT INFORMATION: US 2004038192 A1 20040226

APPLICATION INFO.: US 2003-443452 A1 20030522 (10)

RELATED APPLN. INFO.: Continuation-in-part of Ser. No. US 2001-849618, filed

on 4 May 2001, GRANTED, Pat. No. US 6582953

Continuation-in-part of Ser. No. US 2000-547843, filed

on 12 Apr 2000, GRANTED, Pat. No. US 6642045

NUMBER DATE

PRIORITY INFORMATION: US 1999-129257P 19990414 (60)

DOCUMENT TYPE: Utility FILE SEGMENT: APPLICATION

LEGAL REPRESENTATIVE: HESLIN ROTHENBERG FARLEY & MESITI PC, 5 COLUMBIA

CIRCLE, ALBANY, NY, 12203

NUMBER OF CLAIMS: 10 EXEMPLARY CLAIM: 1

NUMBER OF DRAWINGS: 2 Drawing Page(s)

LINE COUNT: 2256

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

An exsanguinous metabolic support system for maintaining an organ or tissue at a near normal metabolic rate is disclosed that employs a warm perfusion solution capable of altering the production of nitric oxide (NO) in an organ or tissue and supporting the metabolism of the organ or tissue at normothermic temperatures. Perfusion with the solution of the invention, therefore, can be used to regulate nitric oxide production in situations where it is desirable to do so, for example, to prevent reperfusion injury. The system also monitors parameters of the circulating perfusion solution, such as pH, temperature, osmolarity, flow rate, vascular pressure and partial pressure of respiratory gases, and nitric oxide (NO) concentration and regulates them to insure that the organ is maintained under near-physiologic conditions. Use of the system for long-term maintenance of organs for transplantation, for resuscitation and repair of organs having sustained warm ischemic damage, to treat cardiovascular disorders, to prevent reperfusion injury, as a pharmaceutical delivery system and prognosticator of posttransplantation organ function is also disclosed.

IT 56-40-6, Glycine, biological studies 74-79-3, L-Arginine, biological studies 80-68-2, Threonine

83-88-5, Riboflavin, biological studies 302-72-7, Alanine 302-84-1, Serine 516-06-3, Valine

(system for exsanguinous metabolic support of an organ or tissue)

L137 ANSWER 4 OF 33 USPATFULL on STN

ACCESSION NUMBER: 2002:119882 USPATFULL

TITLE: Dosage forms useful for modifying conditions and

functions associated with hearing loss and/or tinnitus

Pearson, Don C., Lakewood, WA, UNITED STATES INVENTOR (S):

Richardson, Kenneth T., Anchorage, AK, UNITED STATES

NUMBER KIND DATE ····· US 2002961870 A1 20020523 US 6524619 B2 20030225 US 2001-765974 A1 20010119 (9) PATENT INFORMATION: APPLICATION INFO.:

> NUMBER DATE -----

PRIORITY INFORMATION:

US 2000-178487P 20000127 (60)

DOCUMENT TYPE:

Utility

FILE SEGMENT:

APPLICATION

LEGAL REPRESENTATIVE: M. Henry Heines, TOWNSEND and TOWNSEND and CREW LLP,

Two Embarcadero Center, 8th Floor, San Francisco, CA,

94111-3834

NUMBER OF CLAIMS: EXEMPLARY CLAIM:

16

LINE COUNT:

2057

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

The invention defines interdependent biofactors and biomolecules, and clinically useful formulations/that are comprised of them. The active agents are demonstrated to be/complementary in their physiologic functions especially as these relate to the quenching of free radicals and to the support of endoth lial physiology, the reduction of hyperinsulinemia and improvements in vascular health. The active components of the invention are selected for inclusion in precise combinations specifically because they improve these various conditions and physiological functions, and by so doing reduce a variety of risks associated with hearing 1ϕ ss and tinnitus. The resulting enhancement of general systemic vascular health, improvement in local VIII.sup.th nerve vascular health, modulation of conditions surrounding blood fluid dynamics, the consequences of hyperinsulinemia, and improvements in free radical defenses, all reduce the potential for cochlear hair cell death and VIII.sup.th nerve atrophy, and the hearing loss and possible deafness that accompany them.

IT 74-79-3, L-Arginine, biological studies 83-88-5, Riboflavin, biological studies 107-35-7, Taurine 107-35-7D, Taurine, reaction with magnesium

> (dosage forms containing vitamin-mineral combinations for modifying conditions and functions associated with hearing loss and tinnitus)

L137 ANSWER 5 OF 33 USPATFULL on STN

ACCESSION NUMBER: 2002:22153 USPATFULL

TITLE: Organ chamber for exsanguinous metabolic support system

INVENTOR(S): Brasile, Lauren, Albany, NY, UNITED STATES

NUMBER KIND DATE US 2002012988 A1 20020131 US 6582953 B2 20030624 PATENT INFORMATION: US 6582953 B2 20030624 US 2001-849618 A1 20010504 (9) APPLICATION INFO.:

Continuation-in-part of Ser. No. US 2000-547843, filed RELATED APPLN. INFO.:

on 12 Apr 2000, PENDING

NUMBER DATE

PRIORITY INFORMATION: WO 2000-US9894 20000413

US 1999-129257P 19990414 (60)

DOCUMENT TYPE: Utility
FILE SEGMENT: APPLICATION

LEGAL REPRESENTATIVE: HESLIN ROTHENBERG FARLEY & MESITI PC, 5 COLUMBIA

CIRCLE, ALBANY, NY, 12203

NUMBER OF CLAIMS: 40 EXEMPLARY CLAIM: 1

NUMBER OF DRAWINGS: 7 Drawing Page(s)

LINE COUNT: 2230

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

An exsanguinous metabolic support system for maintaining an organ or tissue at a near normal metabolic rate is disclosed. The system employs an organ chamber comprising a container and a support member adapted to inhibit movement of the organ within the container during perfusion and/or transport. The organ chamber additionally comprises a conduit for receiving venous outflow of perfusion solution and preventing its contact with the outer surfaces of the organ. A conduit for receiving organ product enables the collection of organ product from a functional organ during perfusion. Use of the organ chamber supports de novo or continued synthesis of constituents necessary for long-term maintenance of organs for transplantation, for resuscitation and active repair of organs that have sustained warm ischemic damage, and for transportation of isolated organs is also disclosed.

IT 56-40-6, Glycine, biological studies 74-79-3, L-Arginine, biological studies 80-68-2, Threonine 83-88-5, Riboflavin, biological studies 302-72-7,

Alanine 302-84-1, Serine 516-06-3, Valine

(system for exsanguinous metabolic support of an organ or tissue)

L137 ANSWER 6 OF 33 USPATFULL on STN

ACCESSION NUMBER: 96:55677 USPATFULL

TITLE: Human liver epithelial cell line and culture media

therefor

INVENTOR(S): Cole, Katharine H., Dayton, MD, United States

Lechner, John F., Bethesda, MD, United States

Reddel, Roger, Camperdown, Australia

Harris, Curtis C., Bethesda, MD, United States

Pfeifer, Andrea M., Pyrbaum, Germany, Federal Republic

of

PATENT ASSIGNEE(S): The United States of America as represented by the Secretary of the Department of Health and Human

Services, Washington, DC, United States (U.S.

government)

NUMBER KIND DATE
US 5529920 19960625
US 1992-879165 19920501 (7)

PATENT INFORMATION:
APPLICATION INFO:

DISCLAIMER DATE: 20120303

RELATED APPLN. INFO.: Continuation-in-part of Ser. No. US 1992-844873, filed on 3 Mar 1992, now patented, Pat. No. US 5342777 which

is a continuation of Ser. No. US 1989-377967, filed on 11 Jul 1989, now abandoned which is a continuation of Ser. No. US 1988-284331, filed on 14 Dec 1988, now

abandoned And a continuation of Ser. No. US 1988-284368, filed on 14 Dec 1988, now abandoned

DOCUMENT TYPE: Utility
FILE SEGMENT: Granted

PRIMARY EXAMINER: Chambers, Jasemine C. ASSISTANT EXAMINER: Stanton, Brian R.

LEGAL REPRESENTATIVE: Birch, Stewart, Kolasch & Birch

NUMBER OF CLAIMS: 2: EXEMPLARY CLAIM: 1

NUMBER OF DRAWINGS: 1 Drawing Figure(s); 1 Drawing Page(s)

LINE COUNT: 1381

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

The present invention relates to long term multiplication and permanent establishment of a cell line of human liver epithelial cells (hepatocytes). The human liver epithelial cell line is capable of mitotically proliferating and continuously growing in vitro under suitable environmental conditions in suitable culture media. A method of producing an immortalized human liver epithelial cell line is also disclosed. The invention also relates to serum-free cell medium developed to support long term multiplication and permanent establishment of a cell line of human liver epithelial cells. The medium may contain an effective cell growth promoting amount of calcium ions; an effective cell growth promoting amount of glucose; an effective amount of insulin to aid cells in glucose uptake; an effective cell growth promoting amount of hydrocortisone; an effective amount of epidermal growth factor to bind epidermal growth factor receptors on cells; an effective amount of transferrin to increase DNA synthesis in cells; an effective amount of cholera toxin to increase DNA synthesis in cells; an effective amount of triiodothyronine to increase DNA synthesis in cells; and an effective growth promoting amount of mammalian hormones

and mitogenic factors, including lipoprotein, cholesterol, phospholipids

and fatty acids.

IT 56-40-6, Glycine, biological studies 56-41-7, Alanine, biological studies 56-45-1, Serine, biological studies 70-26-8, Ornithine 72-18-4, Valine, biological studies 72-19-5, Threonine, biological studies 83-88-5, Riboflavin, biological studies

(human liver epithelial cell line and culture media for it)

L137 ANSWER 7 OF 33 IPA COPYRIGHT 2005 ASHP on STN

ACCESSION NUMBER: 1999:11730 IPA

DOCUMENT NUMBER: 36-12963

TITLE: Stability of TPN solution with or without light shield

package

AUTHOR: Park, K. J.; Park, H. J.; Lee, S. W.; Park, K. H.; Cho, N.

С.

CORPORATE SOURCE: Department of Pharmacy, Seoul National University Hospital,

28, Yongon-Dong, Chongno-Gu, Seoul, 110-744, Korea

SOURCE: ASHP Midyear Clinical Meeting, (Dec 1999) Vol. 34, pp.

INTL-80.

DOCUMENT TYPE: Abstract LANGUAGE: English

ABSTRACT:

We have studied parenteral nutrition solution stability with or without the light shield package. We determined concentrations of each amino acid and vitamins for three days storage at the cool conditions of 2-8DGC. The determined amino acids are isoleucine, leucine, lysine acetate, methionine, phenylalanine, threonine, tryptophan, valine, histidine, ***arginine*** , proline, alanine, glutamic acid, glycine, tyrosine, and serine. The determined vitamins are ascorbic acid, nicotinamide, pyridoxine, thiamine, and riboflavin. We compared TPN solution component stability in plastic bags and glass bottles. The results are as follows: The concentrations of amino acids are all stable in plastic bag or glass bottle for three days. The determined vitamin concentrations are

generally stable but concentrations of ascorbic acid are not stable in glass bottles and plastic bags. The results are glass with light shield (92.2%-89.8%-85.6%), plastic with light shield (93.3%-90.5%-87.4%), glass without light shield(85.7%-90.6%-52.4%) plastic without light shield (79.%-47.6%-30.4%).

SECTION: 10 Drug Stability

INDEX TERM: ASHP meeting abstracts; parenteral nutrition stability

INDEX TERM: Nutrition; parenteral; stability

INDEX TERM: Stability; parenteral nutrition; containers
INDEX TERM: Incompatibilities; parenteral nutrition; storage
INDEX TERM: Containers; parenteral nutrition; incompatibilities

INDEX TERM: Storage; parenteral nutrition; stability

INDEX TERM: Photodecomposition; parenteral nutrition; stability

L137 ANSWER 8 OF 33 BIOSIS COPYRIGHT (c) 2005 The Thomson Corporation. or

STN

ACCESSION NUMBER: 1984:173169 BIOSIS

DOCUMENT NUMBER: PREV198477006153; BA77:6153

TITLE: NUTRITIONAL FACTORS AFFECTING GROWTH AND PRODUCTION OF ANTI

MICROBIAL SUBSTANCES BY STREPTOCOCCUS-LACTIS-SSP-

DIACETYLACTIS S-1-67-C.

AUTHOR(S): REDDY N S [Reprint author]; RANGANATHAN B

CORPORATE SOURCE: DEP ANIM PRODUCTS TECHNOL, FAC AGRIC, OKAYAMA UNIV, OKAYAMA

SHI-700, JPN

SOURCE: Journal of Food Protection, (1983) Vol. 46, No. 6, pp.

514-517.

CODEN: JFPRDR. ISSN: 0362-028X.

DOCUMENT TYPE: Article FILE SEGMENT: BA LANGUAGE: ENGLISH

ABSTRACT: The effect of nutritional factors on the growth and production of antimicrobial substances (AS) by S. lactis ssp. diacetylactis S1-67/C was studied. Among 9 media tested, yeast extract dextrose broth supported good growth and maximum production of AS. Addition of beef extract and yeast extract at 1.0 and 0.6% levels, respectively, increased growth and production of AS. Of 10 carbohydrates examined, maximum production of AS was achieved with 1% glucose followed by fructose, 4% molasses, lactose, sucrose, qalactose, mannitol, maltose and 2% molasses. Xylose inhibited production of AS, although it stimulated growth of the organism. tryptone and tryptose (each at the 1.5% level) significantly stimulated production of AS. Other N sources, including soytone, casein hydrolysate and proteose peptone, retarded production of inhibitory substances. Among the amino acids, L-leucine, DL-methionine and L-glutamic acid were most essential for growth and production of AS, while L-lysine, L-proline, DL-serine , DL-asparatic acid, L-arginine-HCl and DL-tryptophan were stimulatory. Other amino acids such as DL-ornithine, L-cysteine-HCl and DL-citrulline slightly stimulated AS production. In the presence of cynocobalmin, niacin, folic acid, calcium pantothenate and ***riboflavin*** , S. lactis ssp. diacetylactis S1-67/C produced maximum amounts of inhibitory substances. Omission of individual mineral salts from the basal medium did not affect production of AS by the organism. CONCEPT CODE:

Comparative biochemistry 10010
Biochemistry methods - General 10

Biochemistry studies - General 10060 Biochemistry studies - Vitamins 10063

Biochemistry studies - Proteins, peptides and amino acids

10064

Biochemistry studies - Carbohydrates 10068 Biochemistry studies - Minerals 10069

Metabolism - General metabolism and metabolic pathways

13002

Metabolism - Energy and respiratory metabolism 13003

Metabolism - Carbohydrates 13004

Nutrition - General studies, nutritional status and methods

13202

Nutrition - Minerals 13206

Nutrition - Water-soluble vitamins 13210

Nutrition - Carbohydrates 13220

Nutrition - Proteins, peptides and amino acids 13224

Physiology and biochemistry of bacteria 31000 Microbiological apparatus, methods and media 32000 Food microbiology - Food and beverage spoilage and

contamination 39002

Food microbiology - Food and beverage fermentation 39003 Disinfection, disinfectants and sterilization - 39500

Plant physiology - Chemical constituents 51522

INDEX TERMS: Major Concepts

Biochemistry and Molecular Biophysics; Foods;

Metabolism; Nutrition; Physiology

INDEX TERMS: Miscellaneous Descriptors

BEEF EXTRACT YEAST EXTRACT CARBOHYDRATES AMINO-ACIDS

VITAMINS

ORGANISM: Classifier

Gram-Positive Cocci 07700

Super Taxa

Eubacteria; Bacteria; Microorganisms

Taxa Notes

Bacteria, Eubacteria, Microorganisms

ORGANISM: Classifier

Funqi 15000

Super Taxa Plantae Taxa Notes

Fungi, Microorganisms, Nonvascular Plants, Plants

ORGANISM: Classifier

Bovidae 85715

Super Taxa

Artiodactyla; Mammalia; Vertebrata; Chordata; Animalia

Taxa Notes

Animals, Artiodactyls, Chordates, Mammals, Nonhuman

Vertebrates, Nonhuman Mammals, Vertebrates

L137 ANSWER 9 OF 33 BIOSIS COPYRIGHT (c) 2005 The Thomson Corporation. on

STN

ACCESSION NUMBER: 1980:182428 BIOSIS

DOCUMENT NUMBER: PREV198069057424; BA69:57424

TITLE: PRODUCTION OF GENTAMICINS BY MICROMONOSPORA-PURPUREA.

AUTHOR(S): ABOU-ZEID A A [Reprint author]; SALEM H M; EISSA A E-W I

CORPORATE SOURCE: NATL RES CENT, EL-TAHRIR-ST, DOKKI, CAIRO, EGYPT SOURCE: Zentralblatt fuer Bakteriologie Parasitenkunde

Infektionskrankheiten und Hygiene Zweite

Naturwissenschaftliche Abteilung Mikrobiologie der Landwirtschaft der Technologie und des Umweltschutzes,

(1978) Vol. 133, No. 3, pp. 261-275.

DOCUMENT TYPE: Article FILE SEGMENT: BA

LANGUAGE: ENGLISH

ABSTRACT: The natural medium contained the following ingredients (grams per liter): glucose 8.0, or black strap molasses (treated with 0.2-0.3 g/l EDTA) 12.0, fodder yeast (50.0% total nitrogen) 2.0, or fodder yeast (40.0% total nitrogen) 6.0, or yeast extract 8.0, or tryptone 8.0, and

CaCO3 1.0. Treated black strap molasses with EDTA and fodder yeast

proved to be effective in the fermentative production of gentamicins. The most suitable chelating agent was EDTA in the form of disodium for the treatment of

Komombo molasses in a concentration of 0.2-0.3 g/l, while potassium ferrocyanide and methylene blue had depressing effects on the production of gentamicins. The most effective C source, present in Egyptian black strap ***molasses*** , was glucose. Addition of glucose to the medium was preferable at the beginning of the fermentation process. Trace elements present in molasses were very essential for the microbial growth and biosynthesis of gentamicins as proved when molasses ash was added to the natural medium. Organic N sources were more suitable than inorganic N sources for the production of gentamicins by M. purpurea. microorganism utilized the synthetic medium, but the antibiotic yields were less than those produced in the natural medium. The synthetic medium exhibited stimulatory effects of certain amino acids, organic acids, vitamins and purine and pyrimidine bases on the fermentative production of gentamicins. Therefore, the ingredients increasing yields of gentamicins were mainly phenylalanine, isoleucine, lysine, methionine, leucine, arginine, glycine, β-***alanine*** , cystine, tryptophan, malic acid, maleic acid, cobalamin, folic acid, riboflavin, vitamin B1, vitamin B6, biotin, nicotinamide, uracil, adenine, guanine and adenosine. Trace elements (Co, Mo, Fe, Cu, Zn and Mn) exhibited their important role on the biosynthesis and production of

gentamicins by M. purpurea. CONCEPT CODE: Biochem

Biochemistry studies - General 10060

Biochemistry studies - Nucleic acids, purines and

pyrimidines 10062

Biochemistry studies - Vitamins 10063

Biochemistry studies - Proteins, peptides and amino acids

10064

Biochemistry studies - Carbohydrates 10068

Biochemistry studies - Minerals 10069

Metabolism - Carbohydrates 13004

Pharmacology - Drug metabolism and metabolic stimulators

22003

Physiology and biochemistry of bacteria 31000
Microbiological apparatus, methods and media 32000
Chemotherapy - General, methods and metabolism 38502
Food microbiology - Antibiotics, biologics and other agents

39004

Plant physiology - Chemical constituents 51522

Agronomy - Sugar crops 52510

INDEX TERMS:

Major Concepts

Metabolism; Pharmacology; Physiology

INDEX TERMS:

Miscellaneous Descriptors

YEAST EXTRACT GLUCOSE BLACKSTRAP MOLASSES

TRYPTONE ORGANIC ACID VITAMIN PYRIMIDINE TRACE ELEMENTS Classifier

ORGANISM:

Actinoplanetes 08830

Super Taxa

Actinomycetes and Related Organisms; Eubacteria;

Bacteria; Microorganisms

Taxa Notes

Bacteria, Eubacteria, Microorganisms

ORGANISM:

Classifier

Fungi 15000

Super Taxa Plantae Taxa Notes

Fungi, Microorganisms, Nonvascular Plants, Plants

ORGANISM:

Classifier
Angiospermae 25200

Super Taxa

Spermatophyta; Plantae

Taxa Notes

Angiosperms, Plants, Spermatophytes, Vascular Plants

REGISTRY NUMBER: 1403-66-3D (GENTAMICINS)

> 50-99-7Q (GLUCOSE) 58367-01-4Q (GLUCOSE) 289-95-2 (PYRIMIDINE)

8052-35-5 (BLACKSTRAP MOLASSES)

L137 ANSWER 10 OF 33 TOXCENTER COPYRIGHT 2005 ACS on STN DUPLICATE 1

ACCESSION NUMBER: 2000:186295 TOXCENTER

COPYRIGHT:

Copyright 2005 ACS

DOCUMENT NUMBER:

CA13313176534F

TITLE:

Dietetic supplement from human placenta

AUTHOR(S):

Miyares Cao, Carlos Manuel

CORPORATE SOURCE: ASSIGNEE: Centro De Histoterapia Placentaria

PATENT INFORMATION: WO 2000049892 A2 31 Aug 2000 (2000) PCT Int. Appl., 15 pp.

SOURCE:

CODEN: PIXXD2.

COUNTRY:

CUBA Patent CAPLUS

FILE SEGMENT: OTHER SOURCE:

DOCUMENT TYPE:

CAPLUS 2000:608532

LANGUAGE:

Spanish

ENTRY DATE:

Entered STN: 20011116

Last Updated on STN: 20020326

ABSTRACT:

The present invention relates to the field of nutrition, more particularly, to a dietetic supplement obtained from human placenta, which contributes protein and mineral elements to the diet. The tech. aim of the present invention is to provide a new option in the nutritional support to amino acid and mineral salts-deficient patients. The dietetic supplement disclosed has a high nutritional value and shows absolutely no harmful or inconvenient side ***effects*** , thus allowing for its use in both genders and all ages and even during pregnancy. Said material, which is obtained as a residue in the production of medicaments from the human placenta, still contains a considerable amount of highly digestible proteins, vitamins or provitamins and mineral salts that are easily assimilated and is therefore useful as a nutritional supplement in various clin. or surgical conditions characterized by provoking deficient states in individuals suffering from said conditions.

CLASSIFICATION CODE: 17-6

SUPPLEMENTARY TERMS: Miscellaneous Descriptors

dietetic food supplement human placenta

REGISTRY NUMBER:

56-40-6 (Glycine) 56-41-7 (L-Alanine)

56-45-1 (L-Serine)

56-84-8 (L-Aspartic acid)

56-86-0 (L-Glutamic acid)

56-87-1 (L-Lysine)

57-83-0 (Progesterone)

60-18-4 (L-Tyrosine)

61-90-5 (L-Leucine)

63-68-3 (L-Methionine)

63-91-2 (L-Phenylalanine)

64-17-5 (Ethanol)

65-85-0 (Benzoic acid)

67-64-1 (Acetone)

71-00-1 (L-Histidine) 72-18-4 (L-Valine)

72-19-5 (L-Threonine)

74-79-3 (L-Arginine)

83-88-5 (Vitamin B2)

7439-89-6 (Iron)

7440-50-8 (Copper)

7440-66-6 (Zinc) 7440-70-2 (Calcium) 7723-14-0 (Phosphorus)

L137 ANSWER 11 OF 33 TOXCENTER COPYRIGHT 2005 ACS on STN

ACCESSION NUMBER: 2002:280623 TOXCENTER
COPYRIGHT: Copyright 2005 ACS

DOCUMENT NUMBER: CA13801008355H

TITLE: Composition and method for normalizing impaired

or deteriorating neurological function

AUTHOR(S): McCleary, Edward Larry

PATENT INFORMATION: US 2002182196 A1 5 Dec 2002

SOURCE: (2002) U.S. Pat. Appl. Publ., 16 pp.

CODEN: USXXCO.

COUNTRY: UNITED STATES

DOCUMENT TYPE: Patent FILE SEGMENT: CAPLUS

OTHER SOURCE: CAPLUS 2002:928020

LANGUAGE: English

ENTRY DATE: Entered STN: 20021210

Last Updated on STN: 20030624

ABSTRACT:

A nutritional supplement composition for normalizing impaired or deteriorating neurol. function in humans is composed of: at least one agent which promotes synthesis of ATP and/or creatine phosphate in the body, at least one antioxidant for scavenging free radicals in at least one pathway in the body; at least one agent for normalizing or maintaining membrane function and structure in the body; at least one agent for normalizing or maintaining normal neurotransmitter function in the body; at least one agent for down-regulating cortisol action; and at least one agent for suppressing activation of apoptotic pathways in the body. The composition may further contain one or more of: at least one agent for suppressing inflammation in the body; at least one agent for normalizing or maintaining vascular wall function and structure in the body; at least one agent for normalizing or maintaining function of nerve growth factors and/or neurotropic factors in the body; at least one agent for suppressing ***toxic*** metal ionic effects; at least one agent for normalizing or maintaining Me metabolism in the body; at least one agent for normalizing or maintaining metabolism of insulin and glucose in the body; and at least one agent for up-regulating activity of heat shock proteins in the body. A method for normalizing impaired neurol. function in humans modulating nutrient partitioning in a human involves administering the aforementioned composition to the human, preferably on a daily basis, for a therapeutically effective period of Preferably, the method further involves having the human follow a stress reduction program, and/or a cognitive retraining program, and/or a dietary program designed to maximize insulin and glucose metabolism

CLASSIFICATION CODE: 63-6

SUPPLEMENTARY TERMS: Miscellaneous Descriptors

neurol function disorder compn nutraceutical

REGISTRY NUMBER:

50-69-1 (Ribose) 50-81-7 (Vitamin c)

53-43-0 (Dehydroepiandrosterone)

53-84-9 (Nad)
57-00-1 (Creatine)
58-85-5 (Biotin)
59-30-3 (Folic acid)
62-49-7 (Choline)
65-23-6 (Pyridoxine)
68-19-9 (Vitamin b12)
70-51-9 (DESFERRIOXAMINE)
74-79-3 (L-Arginine)
79-83-4 (Pantothenic acid)

83-88-5 (Riboflavin)

98-92-0 (Vitamin b) 107-35-7 (Taurine) 107-43-7 (Betaine) 303-98-0 (Coenzyme q10) 305-84-0 (Carnosine) 501-36-0 (Resveratrol) 502-65-8 (Lycopene) 506-26-3 (γ -Linolenic acid) 987-78-0 (Cytidine 5'-diphosphocholine) 1200-22-2 (α -Lipoic acid) 1406-18-4 (Vitamin e) 3040-38-8 (Acetyl-L-carnitine) 7439-95-4 (Magnesium) 7440-21-3 (Silicon) 7440-66-6 (Zinc) 7782-49-2 (Selenium) 8059-24-3 (Vitamin b6) 12001-76-2 (Vitamin b) 25167-62-8 (Docosahexaenoic acid) 42971-09-5 (Vinpocetine) 58186-27-9 (Idebenone) 102518-79-6 (Huperzine A) 174882-69-0 (Pycnogenol) 56-65-5 (ATP) 67-07-2 (Creatine phosphate) 29908-03-0

REGISTRY NUMBER:

L137 ANSWER 12 OF 33 TOXCENTER COPYRIGHT 2005 ACS on STN

ACCESSION NUMBER: 1999:207222 TOXCENTER COPYRIGHT: Copyright 2005 ACS

DOCUMENT NUMBER:

CA13218236168F

TITLE:

Proximate composition and mineral content of two

edible species of Cnidoscolus (tree spinach)

AUTHOR (S):

Kuti, J. O.; Kuti, H. O.

CORPORATE SOURCE:

College of Agriculture & Human Sciences, Horticultural

Crops Research, Texas A&M University-Kingsville,

Kingsville, TX, 78363, USA.

SOURCE:

Plant Foods for Human Nutrition (Dordrecht, Netherlands),

(1999) Vol. 53, No. 4, pp. 275-283.

CODEN: PFHNE8. ISSN: 0921-9668.

COUNTRY:

UNITED STATES

DOCUMENT TYPE:

Journal CAPLUS

FILE SEGMENT: OTHER SOURCE:

CAPLUS 1999:731679

LANGUAGE:

English

ENTRY DATE:

Entered STN: 20011116

Last Updated on STN: 20020416

ABSTRACT:

Proximate composition and mineral content of raw and cooked leaves of two edible tree spinach species (Cnidoscolus chayamansa and C. aconitifolius), known locally as 'chaya', were determined and compared with that of a traditional green vegetable, spinach (Spinacia oleracea). Results of the study indicated that the edible leafy parts of the two chaya species contained significantly (p < 0.05) greater amts. of crude protein, crude fiber, Ca, K, Fe, ascorbic acid and β -carotene than the spinach leaf. However, no significant (p > 0.05) differences were found in nutritional composition and mineral content between the chaya species, except minor differences in the relative composition of fatty acids, protein and amino acids. Cooking of chaya leaves slightly reduced nutritional composition of both chaya species. Cooking is essential prior to consumption to inactivate the toxic hydrocyanic glycosides present in chaya leaves. Based on the results of this study, the edible chaya leaves may be good dietary sources of minerals (Ca, K and Fe) and vitamins (ascorbic acid and

 β -carotene).

CLASSIFICATION CODE: 17-10

SUPPLEMENTARY TERMS: Miscellaneous Descriptors

Cnidoscolus proximate compn mineral nutrient species;

cooking nutrient chaya leaf

56-41-7 (Alanine) REGISTRY NUMBER:

> 56-85-9 (Glutamine) 56-86-0 (Glutamic acid)

56-87-1 (Lysine)

57-10-3 (Palmitic acid) 57-11-4 (Stearic acid) 60-33-3 (Linoleic acid) 61-90-5 (Leucine) 63-68-3 (Methionine) 63-91-2 (Phenylalanine) 71-00-1 (Histidine)

72-18-4 (Valine) 72-19-5 (Threonine) 73-32-5 (Isoleucine) 74-79-3 (Arginine)

112-80-1 (Oleic acid) 463-40-1 (Linolenic acid) 59-43-8 (Thiamin)

83-88-5 (Riboflavin) 7440-23-5 (Sodium) 7439-95-4 (Magnesium) 50-81-7 (Ascorbic acid) 7235-40-7 (β -Carotene)

7439-89-6 (Iron) 7440-09-7 (Potassium) 7440-70-2 (Calcium)

L137 ANSWER 13 OF 33 TOXCENTER COPYRIGHT 2005 ACS on STN

1999:140598 TOXCENTER ACCESSION NUMBER: COPYRIGHT:

Copyright 2005 ACS

DOCUMENT NUMBER:

CA13106072895S

TITLE:

Chemical composition and biological evaluation

of mahua flowers

AUTHOR (S):

Jayasree, B.; Harishankar, N.; Rukmini, C.

CORPORATE SOURCE:

National Institute of Nutrition, Indian Council of Medical

Research, Hyderabad, 500007, India.

SOURCE:

Journal of the Oil Technologists' Association of India (Mumbai, India), (1998) Vol. 30, No. 4, pp. 170-172.

CODEN: JOTIAC. ISSN: 0970-4094.

COUNTRY:

TNDTA DOCUMENT TYPE: Journal FILE SEGMENT: CAPLUS

OTHER SOURCE:

CAPLUS 1999:258372

LANGUAGE:

English

ENTRY DATE:

Entered STN: 20011116

Last Updated on STN: 20020416

ABSTRACT:

Mahua (Madhuca latifolia, Sapotaceae) flowers were defatted and desugared and analyzed for their nutrient composition and the protein quality evaluated in weanling rats. The flowers are a good source of sugars (68%), calcium, phosphorus and protein (6.67%). Lysine content of the flower protein is higher than any cereal protein and also a good source of sulfur containing amino acids. Its PER and NPU were comparable to those of control group. It did not show any ***toxic*** symptoms. Mahua flowers may form a good dietary source for

tribals.

CLASSIFICATION CODE: 17-10

SUPPLEMENTARY TERMS: Miscellaneous Descriptors

```
mahua flower nutrient protein quality
REGISTRY NUMBER:
                     57-10-3 (Palmitic acid)
                     57-11-4 (Octadecanoic acid)
                     60-33-3 (9,12-Octadecadienoic acid (9Z,12Z)-)
                     112-80-1 (9-Octadecenoic acid (9Z)-)
                     112-85-6 (Behenic acid)
                     124-07-2 (Octanoic acid)
                     142-62-1 (Caproic acid)
                     143-07-7 (Lauric acid)
                     334-48-5 (Capric acid)
                     373-49-9 (Palmitoleic acid)
                     463-40-1 (Linolenic acid)
                     544-63-8 (Myristic acid)
                     544-64-9 (Myristoleic acid)
                     557-59-5 (Lignoceric acid)
                     22032-47-9 (Lauroleic acid)
                     50-81-7 (L-Ascorbic acid)
                       56-40-6 (Glycine)
                       56-41-7 (L-Alanine)
                       56-45-1 (L-Serine)
                     56-84-8 (L-Aspartic acid)
                     56-86-0 (L-Glutamic acid)
                     56-87-1 (L-Lysine)
                     56-89-3 (L-Cystine)
                     59-43-8 (Thiamine)
                     59-67-6 (Niacin)
                     60-18-4 (L-Tyrosine)
                     61-90-5 (L-Leucine)
                     63-68-3 (L-Methionine)
                     63-91-2 (L-Phenylalanine)
                     71-00-1 (L-Histidine)
                       72-18-4 (L-Valine)
                       72-19-5 (L-Threonine)
                     73-32-5 (L-Isoleucine)
                       74-79-3 (L-Arginine)
                       83-88-5 (Riboflavin)
                     147-85-3 (L-Proline)
                     7440-70-2 (Calcium)
                     7723-14-0 (Phosphorus)
L137 ANSWER 14 OF 33 TOXCENTER COPYRIGHT 2005 ACS on STN
ACCESSION NUMBER:
                     1998:128554 TOXCENTER
COPYRIGHT:
                     Copyright 2005 ACS
DOCUMENT NUMBER:
                     CA12822269798G
                     Study on the chemical change of amino acid and
TITLE:
                     vitamin of rapeseed during germination
AUTHOR (S):
                     Kim, In-Sook; Han, Sung-Hee; Han, Kwang-Wan
                     Dep. Food and Nutrition, Wonkwang Univ., Cheonbuk,
CORPORATE SOURCE:
                     570-749, S. Korea.
SOURCE:
                     Han'guk Sikp'um Yongyang Kwahak Hoechi, (1997) Vol. 26,
                     No. 6, pp. 1058-1062.
                     CODEN: HSYHFB. ISSN: 1226-3311.
                     KOREA, REPUBLIC OF
COUNTRY:
DOCUMENT TYPE:
                     Journal
FILE SEGMENT:
                     -CAPLUS
                     CAPLUS 1998:217893
OTHER SOURCE:
LANGUAGE:
                     Korean
                     Entered STN: 20011116
ENTRY DATE:
                     Last Updated on STN: 20020605
ABSTRACT:
The objective of this was to investigate the tech. feasibility of producing
```

toxicant -free by germination. To this end, rapeseed (Brassica napus

L.) was germinated at 25°C for 120 h, and the chemical compns. of amino acids and vitamins were determined every 24 h during germination. germination, rapeseed contained 5.4 g/16 g N of glutamic acid and high percentage of the other amino acids in order of Asp > Leu > His > Pro > Arg > Lys > Gly > Ser > Ala > Val. The amino acids were gradually decreased until 96 h during germination had tendency to show a slight increase in 120 h. Vitamin B1, B2 and C contents in rapeseed before germination were 0.11, 0.21 and 3.72 mg% resp., and the vitamin E was 423 $\mu g/g$. The vitamin C greatly increased in 72 h during germination, while the vitamin B group was drastically decreased in 72 h. Thus, germination process is very effective to the removal of ***toxicants*** in rapeseed.

CLASSIFICATION CODE: 17-10

SUPPLEMENTARY TERMS: Miscellaneous Descriptors

rapeseed germination amino acid vitamin change

REGISTRY NUMBER:

50-81-7 (Vitamin C) 52-90-4 (L-Cysteine) 56-40-6 (Glycine) 56-41-7 (L-Alanine) 56-45-1 (L-Serine) 56-84-8 (L-Aspartic acid)

56-86-0 (L-Glutamic acid)

56-87-1 (L-Lysine) 59-02-9 (α -Tocopherol) 59-43-8 (Vitamin B1) 60-18-4 (L-Tyrosine) 61-90-5 (L-Leucine) 63-68-3 (L-Methionine) 63-91-2 (L-Phenylalanine) 71-00-1 (L-Histidine) 72-18-4 (L-Valine) 72-19-5 (L-Threonine)

73-22-3 (L-Tryptophan) 73-32-5 (L-Isoleucine) 74-79-3 (L-Arginine) 83-88-5 (Vitamin B2) 147-85-3 (L-Proline) 1406-18-4 (Vitamin E)

7616-22-0 (γ -Tocopherol)

L137 ANSWER 15 OF 33 TOXCENTER COPYRIGHT 2005 ACS on STN

ACCESSION NUMBER: COPYRIGHT:

1993:150746 TOXCENTER Copyright 2005 ACS

DOCUMENT NUMBER:

CA11902014946A

TITLE:

Bioaccumulation of toxicants, element and

nutrient composition, and soft tissue histology of zebra mussels (Dreissena polymorpha) from New York

State waters

AUTHOR (S):

Secor, Carol L.; Mills, Edward L.; Harshbarger, John; Kuntz, H. Thomas; Gutenmann, Walter H.; Lisk, Donald J.

CORPORATE SOURCE:

Cornell Biol. Field Stn., Bridgeport, NY, 13030, USA. Chemosphere, (1993) Vol. 26, No. 8, pp. 1559-75.

CODEN: CMSHAF. ISSN: 0045-6535.

COUNTRY:

SOURCE:

UNITED STATES

DOCUMENT TYPE: FILE SEGMENT:

Journal CAPLUS

OTHER SOURCE:

CAPLUS 1993:414946

LANGUAGE:

English

ENTRY DATE:

Entered STN: 20011116

Last Updated on STN: 20020917

ABSTRACT:

Zebra mussels (Dreissena polymorpha) were collected from 3 rivers and 3 lakes in New York State and analyzed for toxic and nutrient elements, amino

acids, and vitamins. The concentration of Cd and Se in soft tissues was generally Ca comprised 40% by weight of the shell. Polychlorinated biphenyls were markedly higher in soft tissues of zebra mussels from the Hudson River than the other waters. Mussel soft tissues from only 2 waters showed detectable levels of p,p'-DDE. Significant histol. lesions or infectious agents were not observed in soft tissues. CLASSIFICATION CODE: 61-2 SUPPLEMENTARY TERMS: Miscellaneous Descriptors Dreissena polymorpha toxic compd bioaccumulation; river water pollution Dreissena toxic bioaccumulation; lake water pollution Dreissena toxic bioaccumulation REGISTRY NUMBER: 56-40-6 (Glycine) 56-41-7 (Alanine) 56-45-1 (Serine) 56-84-8 (Aspartic acid) 56-86-0 (Glutamic acid) 56-87-1 (Lysine) 56-89-3 (Cystine) 59-43-8 (Vitamin b1) 59-67-6 (Niacin) 60-18-4 (Tyrosine) 61-90-5 (Leucine) 63-68-3 (Methionine) 63-91-2 (Phenylalanine) 71-00-1 (Histidine) 72-18-4 (Valine) 72-19-5 (Threonine) 73-32-5 (Isoleucine) 74-79-3 (Arginine) 83-88-5 (Vitamin b2) 147-85-3 (Proline) 7439-89-6 (Iron) 7439-92-1 (Lead) 7439-95-4 (Magnesium) 7439-96-5 (Manganese) 7439-97-6 (Mercury) 7439-98-7 (Molybdenum) 7440-02-0 (Nickel) 7440-09-7 (Potassium) 7440-23-5 (Sodium) 7440-42-8 (Boron) 7440-43-9 (Cadmium) 7440-47-3 (Chromium) 7440-50-8 (Copper) 7440-62-2 (Vanadium) 7440-66-6 (Zinc) 7440-70-2 (Calcium) 7704-34-9 (Sulfur) 7723-14-0 (Phosphorus) 7727-37-9 (Nitrogen) 7782-49-2 (Selenium) 11097-69-1 (Aroclor 1254) 11103-57-4 (Vitamin a)

REGISTRY NUMBER:

72-55-9

L137 ANSWER 16 OF 33 TOXCENTER COPYRIGHT 2005 ACS on STN

ACCESSION NUMBER: COPYRIGHT:

1952:2946 TOXCENTER Copyright 2005 ACS

12672-29-6 (Aroclor 1248)

DOCUMENT NUMBER:

CA04617049681M

TITLE:

Effect of B vitamins and amino acids on the rate

of multiplication of Paramecium caudatum

AUTHOR(S):

Kreitmaier, Georg

CORPORATE SOURCE:

Zool. Inst. Munich, Germany.

SOURCE:

Zeitschrift fuer Vitamin-, Hormon- und Fermentforschung,

(1952) Vol. 4, pp. 542-54.

CODEN: ZVHFAW. ISSN: 0373-0220.

COUNTRY:

GERMANY, FEDERAL REPUBLIC OF

DOCUMENT TYPE:

Journal **CAPLUS**

FILE SEGMENT: OTHER SOURCE:

CAPLUS 1952:49681

LANGUAGE:

German

ENTRY DATE:

Entered STN: 20011116

Last Updated on STN: 20030902

ABSTRACT:

The effect of adding these supplements, singly and in combination, to the basal Knop medium was studied: thiamine, riboflavin, pyridoxine, nicotinamide, folic acid, d-biotin, p-aminobenzoic acid, pantothenic acid, vitamin B12, inositol, choline, arginine, asparagine, β - ***alanine*** , cysteine, glutamic acid, histidine, and glutathione. Natural materials (casein, yeast extract, wheat germ, mammalian liver and heart exts.,

etc.) were used for comparison. The rate of multiplication is accelerated by optimal levels of these supplements (usually 0.1-10 γ/ml.), while higher levels usually are toxic.

CLASSIFICATION CODE: 11I

REGISTRY NUMBER:

150-13-0 (Benzoic acid, p-amino-)

68-19-9 (Vitamin, B12) 56-86-0 (Glutamic acid) 59-30-3 (Folic acid) 59-43-8 (Vitamin, B1) 70-18-8 (Glutathione) 71-00-1 (Histidine) 83-88-5 (Vitamin, B2) 8059-24-3 (Vitamin, B6)

58-85-5 (Biotin) 62-49-7 (Choline) 74-79-3 (Arginine) 87-89-8 (Inositol) 107-95-9 (β - Alanine) 52-90-4 (Cysteine) 70-47-3 (Asparagine) 98-92-0 (Nicotinamide)

79-83-4 (Pantothenic acid)

L137 ANSWER 17 OF 33 TOXCENTER COPYRIGHT 2005 ACS on STN

ACCESSION NUMBER:

1954:3886 TOXCENTER

COPYRIGHT:

Copyright 2005 ACS

DOCUMENT NUMBER:

CA04808025941M

TITLE:

Amino acids and growth factors in chemically

defined medium for Drosophila

AUTHOR(S):

Hinton, T.; Noyes, D. T.; Ellis, J.

SOURCE:

Physiological Zoology, (1951) Vol. 24, pp. 335-53.

CODEN: PHZOA9. ISSN: 0031-935X.

DOCUMENT TYPE:

Journal

FILE SEGMENT: OTHER SOURCE: CAPLUS

ENTRY DATE:

CAPLUS 1954:25941 Entered STN: 20011116

Last Updated on STN: 20040316

ABSTRACT:

Drosophila does not utilize D-tryptophan, while high concns. of the L-form are ***toxic*** ; high concns. of L-isoleucine and L-serine slightly

inhibit growth, while D-serine is extremely toxic. The

arginine requirement of Drosophila can be met partly by

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***citrulline.***
                      Glycine is essential for optimal growth and also
exerts a detoxifying effect on other essential amino acids. Biotin, which is
spared by citrulline in the absence of arginine, can be
tolerated in high concentration; it cannot be replaced by lecithin. High concns. of
pyridoxine inhibited development while vitamin B12 slightly increased the
percent of larvae pupating. Protogen had no effect on development, while
vitamin B12 improved pupation but inhibited development, both only slightly.
Inositol, p-aminobenzoic acid, and various yeast fractions had no effect on
development. The following medium, completely defined except for the agar
base, allows almost normal development (in mg./ml.): alanine 1.085,
***arginine***
                 0.794, aspartic acid 1.221, cystine 0.480, glutamic acid
4.418, glycine 1.745, histidine 0.484, hydroxyproline 0.384,
isoleucine 1.260, leucine 2.345, lysine 1.337, methionine 0.339, phenylalanine
1.008, proline 1.682, threonine 0.758, tryptophan 1.745, tyrosine
1.240, valine 1.335, sucrose 7.5, cholesterol 0.1, ergosterol 1.0, ribonucleic acid 1.0, inosine 0.25, thymine 0.004, biotin 0.00002, vitamin B12
0.00004, Ca pantothenate 0.006, choline chloride 0.020, pteroylglutamic acid
0.006, pyridoxine 0.030, riboflavine 0.0024, thiamine 0.0015, and nicotinamide
0.010. Tatum's salt mixture was also included.
CLASSIFICATION CODE: 11I
                      68-19-9 (Vitamin, B12)
REGISTRY NUMBER:
                        56-45-1 (Serine)
                      73-22-3 (Tryptophan)
                      73-32-5 (Isoleucine)
                      8059-24-3 (Vitamin, B6)
                        372-75-8 (Citrulline)
                      51-35-4 (Proline, hydroxy-)
                        56-40-6 (Glycine)
                        56-41-7 (Alanine)
                      56-84-8 (Aspartic acid)
                      56-86-0 (Glutamic acid)
                      56-87-1 (Lysine)
                      56-89-3 (Cystine)
                      57-50-1 (Sucrose)
                      57-87-4 (Ergosterol)
                      57-88-5 (Cholesterol)
                      58-85-5 (Biotin)
                      59-30-3 (Folic acid)
                      59-43-8 (Vitamin, B1)
                      60-18-4 (Tyrosine)
                      61-90-5 (Leucine)
                      62-49-7 (Choline)
                      63-68-3 (Methionine)
                      63-91-2 (Alanine, phenyl-)
                      65-71-4 (Thymine)
                      71-00-1 (Histidine)
                        72-18-4 (Valine)
                        72-19-5 (Threonine)
                        74-79-3 (Arginine)
                        83-88-5 (Vitamin, B2)
                      86-04-4 (Inosine, diphosphate)
                      98-92-0 (Nicotinamide)
                      137-08-6 (Pantothenic acid, calcium salt)
                      147-85-3 (Proline)
L137 ANSWER 18 OF 33 TOXCENTER COPYRIGHT 2005 ACS on STN
ACCESSION NUMBER:
                      2001:293113 TOXCENTER
COPYRIGHT:
                      Copyright 2005 ACS
DOCUMENT NUMBER:
                      CA03814025518X
TITLE:
                      Amino acid mixtures effective parenterally for
                      long-continued plasma protein production. Casein digests
```

compared

AUTHOR(S): Madden, S. C.; Woods, R. R.; Shull, F. W.; Whipple, G. H. SOURCE: Journal of Experimental Medicine, (1944) Vol. 79, pp.

607-24.

CODEN: JEMEAV. ISSN: 0022-1007.

DOCUMENT TYPE: Journal FILE SEGMENT: CAPLUS

OTHER SOURCE: CAPLUS 1944:25518
ENTRY DATE: Entered STN: 20011218

Last Updated on STN: 20030624

ABSTRACT:

When blood plasma proteins are depleted by bleeding with return of red cells suspended in saline (plasmapheresis), it is possible to bring dogs to a steady state of hypoproteinemia and a constant level of plasma protein production if the diet N intake is controlled and limited. Such dogs are outwardly normal but have a lowered resistance to infection and to certain intoxications. The 10 growth-essential amino acids of Rose plus glycine will maintain N balance and produce as much new plasma protein as will good diet proteins. This good utilization is demonstrated over periods of several months when the amino acids are given either orally or parenterally. There is no evidence of ***toxicity*** in general nor to unnatural forms of these synthetic amino acids in particular. Given parenterally appropriate mixts. of these amino acids are well tolerated even upon rapid injection. The min. daily requirements for a 10-kg. dog can be given intravenously in 10 min. without reaction. Subcutaneously, a 10% solution can be given rapidly without reaction. Among various mixts. tested, the following approximates a min. for a 10-kg. dog: dl-threonine 0.7, dl-valine 1.5, l(-)-leucine 1.5, dl-isoleucine 1.4, dl-lysine-HCl 0.5, dl-phenylalanine 1, l-(-)-tryptophan 0.4, dl-methionine 0.6, 1(+)-histidine-HCl 0.5, 1(+)-arginine-HCl 0.5 and ***glycine*** 1 g. The presence of glycine improves tolerance to rapid intravenous injection but excess glycine does not improve utilization of the mixture Over long periods, this mixture appears sub-optimal in quantity; doubled, it is more ample. Of 2 casein digests tested, the one prepared by enzymic hydrolysis provided good N retention and fairly good plasma protein production but was much less tolerable upon intravenous injection than certain mixts. of pure amino acids. The one prepared by acid hydrolysis and tryptophan fortification afforded bare N equilibrium and produced virtually no plasma protein. Skin lesions observed after 10-20 weeks of synthetic diet probably reflect a deficiency of some member or members of the vitamin ***B2*** group. A persistent slight weight loss in the face of a strongly pos. N balance may accompany this deficiency. CLASSIFICATION CODE: 11H

L137 ANSWER 19 OF 33 WPIDS COPYRIGHT 2005 THE THOMSON CORP on STN

ACCESSION NUMBER: 2004-267967 [25] WPIDS

CROSS REFERENCE: 2003-102288 [09] DOC. NO. CPI: C2004-104361

TITLE: Metabolic uncoupling therapy involves formulating a

combination of agent of metabolic uncoupling therapy that

limits the accumulation of high-energy electrons

potentially available to the electron transport chain.

DERWENT CLASS: B05

INVENTOR(S): MCCLEARY, E L

PATENT ASSIGNEE(S): (MCCL-I) MCCLEARY E L

COUNTRY COUNT: 1

PATENT INFORMATION:

APPLICATION DETAILS:

PATENT NO	KIND	APPLICATION	DATE
US 2004043013	_	US 2000-749584 US 2003-462958	20001228

FILING DETAILS:

PATENT NO KIND PATENT NO US 2004043013 A1 CIP of US 6579866

PRIORITY APPLN. INFO: US 2003-462958 200300 2000-749584 20001228 20030617; US

INT. PATENT CLASSIF.:

MAIN: A61K031-7076 NDARY: A61K031-195; A61K031-198; A61K031-525; A61K031-685 SECONDARY:

BASIC ABSTRACT:

US2004043013 A UPAB: 20040418

NOVELTY - Metabolic uncoupling therapy (MUT) involves analyzing specific physiologic process, including delineating the metabolic pathways related to reductive stress; identifying several MUT agents that modulate the metabolic pathways by influencing electron flux; and formulating combination of MUT agent that limits the accumulation of high-energy electrons potentially available to the electron transport chain.

DETAILED DESCRIPTION - An INDEPENDENT CLAIM is included for a composition comprising at least two (preferably at least three, especially at least five) small electrophilic biomolecules (a), at least one oxaloacetate precursor (b), at least two vitamin B and its structurally related entity (c) (preferably at least three, especially at least five); and at least one electron cycling agent (d). (a) Is TMG (trimethylglycine), choline, phosphatidyl choline, SAMe (S-adenosyl methionine), carnitine, ALC (acetyl L-carnitine), propionyl carnitine, myoinositol, sphingomyelin, glycerylphosphorylcholine or acetylcholine. (b) Is pyruvate, aspartate, glycine or serine. (c) Is folate, riboflavin, B1, B3, niacinamide, nicotinamide, polynicotinate, B6, B12, biotin, pantothenic acid, riboflavin or related chemical species. (d) Is coenzyme Q10, lipoic acid or

ACTIVITY - Antidiabetic; Antilipemic; Antiinflammatory; Vasotropic; Cardiant; Cerebroprotective; Anorectic; Nootropic; Tranquilizer; Muscular-Gen.; Dermatological; Neuroprotective; Gastrointestinal-Gen.; Hepatotropic; Virucide. Test details are given, but no results are given. MECHANISM OF ACTION - None given.

USE - For metabolite uncoupling therapy (claimed), which is useful for the prevention of a multitude of conditions and as a therapeutic modality under conditions of disease e.g. high blood pressure, diabetes, dyslipidemia, hyperlipidemia, hypercholesterolemia, insulin resistance, inflammation, vascular disease, heart disease, stroke, overweight, obesity, neuronal and/or cognitive dysfunction, dementia, attention and attention/hyperactivity disorder, mood disorder, muscular damage, muscular deterioration or soreness, athletic compromise, sarcopenia, glucose intolerance and other disorders of glucose metabolism, premature aging, skin deterioration and/or damage either associated with, or not associated with sun exposure, loss of muscle tone, frailty and bone loss, and aging. The composition is useful in food products e.g. milk or milk products, juices, shakes, salad dressing, gravies, sauces, nutritional bars, protein powders and any other palatable food products and in enhancement of athletic performance in greyhounds or racehorses, enhanced and prolonged fertility in breeding stock and health maintenance in household pets and as a brain performance-enhancing drink mix. For the treatment of neurodegenerative disorders e.g. multiple sclerosis and Alzheimer's disease, inflammatory gastrointestinal disorder and hepatic

steatosis/steatohepatitis.

ADVANTAGE - The combination of the MUT agents limits the accumulation of high-energy electrons potentially available to the electron transport chain. The MUT includes manipulation of flux of high-energy electrons through biochemical pathways; modulation of related cell processes and signaling systems, modulation of metabolic intermediates involved in the production of high energy electrons and modulation of nucleotides, nucleotide ratios and nucleotide cycling. The MUT minimizes adverse side effects that might occur through

inappropriate usage of various compound and composition not in accordance with the combination of the MUT agents.

Dwg.0/0

FILE SEGMENT: CPI FIELD AVAILABILITY: AB; DCN

MANUAL CODES: CPI: B03-B; B03-C; B03-D; B03-E; B03-G; B04-A08C2;

B04-A10B; B04-B03A; B04-B04D2; B04-J03A; B05-A01A; B05-A01B; B05-A03A; B05-A03B; B05-B01M; B05-B01P; B07-H; B10-A06; B10-A09B; B10-A17; B10-A22; B10-B02; B10-C04B; B10-C04E; B11-C08E; B12-K04A; B14-C03; B14-D02A2; B14-E10; B14-E12; B14-F01B; B14-F02B; B14-F02F; B14-F06; B14-J01; B14-J05; B14-N01;

B14-N12; B14-N16; B14-N17; B14-S01

L137 ANSWER 20 OF 33 WPIDS COPYRIGHT 2005 THE THOMSON CORP on STN

ACCESSION NUMBER: 2004-464569 [44] WPIDS

DOC. NO. CPI: C2004-173882

TITLE: Health food/pharmaceutical, useful for preventing e.g.

diabetes, comprises zinc complex having ligand containing

amino acids, picolinic acids, vitamins, maltols,

carboxylic acids, oligopeptides or their derivatives, and

zinc source.

DERWENT CLASS: B05 D13

PATENT ASSIGNEE(S): (ARIT-I) ARITA J

COUNTRY COUNT: 1

PATENT INFORMATION:

PATENT NO KIND DATE WEEK LA PG MAIN IPC

JP 2004175790 A 20040624 (200444)* 8 A61K031-315

APPLICATION DETAILS:

PATENT NO KIND APPLICATION DATE
-----JP 2004175790 A JP 2003-374551 20031104

PRIORITY APPLN. INFO: JP 2002-327684 20021112

INT. PATENT CLASSIF.:

MAIN: A61K031-315

SECONDARY: A23L001-304; A61K031-19; A61K031-191; A61K031-192;

A61K031-194; A61K031-198; A61K031-205; A61K031-351; A61K031-375; A61K031-401; A61K031-405; A61K031-4172; A61K031-4402; A61K031-455; A61K031-519; A61K031-525;

A61K031-555; A61K038-00; A61P003-10; A61P043-00

BASIC ABSTRACT:

JP2004175790 A UPAB: 20040712

NOVELTY - A health food/pharmaceutical comprises zinc complex having ligand containing amino acids, picolinic acids, vitamins, maltols, carboxylic acids, oligopeptides or their derivatives complexed with zinc source.

ACTIVITY - Antidiabetic.

MECHANISM OF ACTION - Glucosidase-Inhibitor-Alpha.

(alpha)-glucosidase inhibiting effect of zinc complex (Zn(Gln)2) was performed by improving the method described in Japanese Patent No.2002316939. The (alpha)-glucosidase inhibiting effect was 5.4 micro

USE - The health food/pharmaceutical is useful for preventing life-style diseases e.g. diabetes and hyperglycemia.

ADVANTAGE - The health food/pharmaceutical having excellent (alpha)-glucosidase inhibiting effect can be administered safely for prolonged period. The health food effectively promotes carbohydrate metabolism without causing any side effect.

Dwg.0/7

FILE SEGMENT: CPI FIELD AVAILABILITY: AB; DCN

MANUAL CODES:

CPI: B03-B; B03-C; B03-F; B03-L; B04-C01; B05-A03A; B05-C07; B06-D01; B06-D09; B06-D17; B07-A03; B07-D03; B07-D04C; B07-D09; B07-D12; B07-F01; B10-A07; B10-A17; B10-A22; B10-B01B; B10-B02D; B10-B02E; B10-B02H; B10-B02J; B10-C02; B10-C03; B10-C04D; B10-C04E; B14-D07B; B14-E11; B14-F09; B14-S04; D03-H01T2

L137 ANSWER 21 OF 33 WPIDS COPYRIGHT 2005 THE THOMSON CORP on STN

ACCESSION NUMBER:

2004-456017 [43] WPIDS

DOC. NO. CPI:

C2004-170976

TITLE:

Foodstuff e.g. nutritional food with antioxidant effect for improving lifestyle disease e.g. diabetes, comprises ligand forming zinc source, organic compound containing basic amino acid for forming complex with zinc and

vitamin.

DERWENT CLASS:

B05 D13

PATENT ASSIGNEE(S):

(ARIT-I) ARITA J

COUNTRY COUNT:

PATENT INFORMATION:

PATENT NO KIND DATE WEEK LA PG MAIN IPC ______ JP 2004166690 A 20040617 (200443)* 6 A23L001-304

APPLICATION DETAILS:

APPLICATION DATE PATENT NO KIND _____ JP 2003-363204 20031023 JP 2004166690 A

PRIORITY APPLN. INFO: JP 2002-319193

20021101

INT. PATENT CLASSIF.:

MAIN: A23L001-304

SECONDARY:

A23L001-302; A61K033-30; A61P039-06

BASIC ABSTRACT:

JP2004166690 A UPAB: 20040709

NOVELTY - Foodstuff comprises a source of zinc which forms a ligand, an organic compound containing basic amino acid except histidine, which form a complex with zinc, and a vitamin or its derivatives.

ACTIVITY - Antidiabetic; Antiarteriosclerotic; Cytostatic; Dermatological.

MECHANISM OF ACTION - None given.

USE - Used as nutritional food, health food and health supplement having antioxidant effect, for improving lifestyle related disease e.g. diabetes, arteriosclerosis, cancer and aging caused by active oxygen and as antioxidant in another foodstuff, food additive, vitamins and/or

minerals.

In a test, the influence of zinc/vitamin C complex having antioxidant action with respect to acute renal failure by cisplatin involving active oxygen was evaluated. 7.5 mg/kg of cisplatin was administered intravenously to male Sprague Dawley rats. Urine and blood samples were collected after 4-18 days of administration. 100 mg/kg of zinc/vitamin C complex was dissolved in distilled water and administered orally to the rats, once daily. Cisplatin increased blood urea nitrogen and N-acetyl-(beta)-D-glucosaminidase. The zinc/vitamin C prevented renal disease caused by cisplatin by reducing active oxygen.

ADVANTAGE - Zinc (II) complex having a ligand vitamin such as vitamin C has reduced toxicity, good degree of stability and lipophilicity.

Dwg.0/4

FILE SEGMENT: CPI FIELD AVAILABILITY: AB; DCN

MANUAL CODES: CPI: B03-L; B05-A03A; B10-B02C; B14-E11; B14-F07;

B14-H01; B14-N17; B14-S04; B14-S08; D03-H01T2

L137 ANSWER 22 OF 33 WPIDS COPYRIGHT 2005 THE THOMSON CORP on STN

ACCESSION NUMBER: 2003-221666 [21] WPIDS

DOC. NO. CPI: C2003-056433

TITLE: Composition used in prophylaxis and/or treatment of

symptoms caused or exacerbated by consumption of toxic compound such as ethanol comprises fructose

and/or fructose containing oligosaccharide.

DERWENT CLASS: B05

INVENTOR(S): MCGREGOR, N R

PATENT ASSIGNEE(S): (PENA-N) PENAM INVESTMENTS PTY LTD; (MCGR-I) MCGREGOR N R

COUNTRY COUNT: 101

PATENT INFORMATION:

PATENT NO KIND DATE WEEK LA PG MAIN IPC

WO 2003006073 A1 20030123 (200321)* EN 12 A61K031-7004

RW: AT BE BG CH CY CZ DE DK EA EE ES FI FR GB GH GM GR IE IT KE LS LU

MC MW MZ NL OA PT SD SE SK SL SZ TR TZ UG ZM ZW

W: AE AG AL AM AT AU AZ BA BB BG BR BY BZ CA CH CN CO CR CU CZ DE DK DM DZ EC EE ES FI GB GD GE GH GM HR HU ID IL IN IS JP KE KG KP KR KZ LC LK LR LS LT LU LV MA MD MG MK MN MW MX MZ NO NZ OM PH PL PT RO RU SD SE SG SI SK SL TJ TM TN TR TT TZ UA UG US UZ VN YU ZA ZM

ZW 879 A1 20040428 (200429) EN A61K006-00

EP 1411879 A1 20040428 (200429) EN A61K006-00 R: AL AT BE BG CH CY CZ DE DK EE ES FI FR GB GR IE IT LI LT LU LV MC

MK NL PT RO SE SI SK TR

AU 2002318969 A1 20030129 (200452) A61K031-7004 JP 2004534094 W 20041111 (200474) 36 A61K031-7004 US 2004248819 A1 20041209 (200481) A61K031-70

APPLICATION DETAILS:

PATENT NO	KIND	APPLICATION	DATE
WO 2003006073 EP 1411879	A1 A1	WO 2002-AU890 EP 2002-748426	20020705
		WO 2002-AU890	20020705
AU 2002318969 JP 2004534094	A1 W	AU 2002-318969 WO 2002-AU890	20020705 20020705
US 2004248819	A1	JP 2003-511878 WO 2002-AU890	20020705 20020705
		US 2004-483393	20040628

FILING DETAILS:

PATENT NO	KIND	PATENT NO					
EP 1411879	Al Based on	WO 2003006073					
AU 2002318969	Al Based on	WO 2003006073					
JP 2004534094	W Based on	WO 2003006073					

PRIORITY APPLN. INFO: AU 2001-6261 20010710

INT. PATENT CLASSIF.:

MAIN: A61K006-00; A61K031-70; A61K031-7004

SECONDARY: A61K031-197; A61K031-198; A61K031-381; A61K031-385; A61K031-4188; A61K031-455; A61K031-51; A61K031-525;

A61K031-7016; A61K031-702; A61K031-715; A61P025-32;

A61P039-02

BASIC ABSTRACT:

WO2003006073 A UPAB: 20030328

NOVELTY - Composition comprises fructose and/or a fructose-containing oligosaccharide.

ACTIVITY - Antialcoholic.

In a test, a male subject aged 42 years, who is a moderate drinker, took a teaspoon of a supplement before and after drinking. The supplement comprised (in mg per 100 g) corn maltodextrins (56609.1), fructose (21000), dextrose monohydrate (7000), L-alanine (3500), L-leucine (2500), L-isoleucine (2500), L-valine (2500), Lglycine (1000), L-serine (500), L-methionine (50), L-phenylalanine (50), L-arginine (50), L-tyrosine (50), L-histidine (50), L-aspartic acid (50), L-glutamic acid (50), L-asparagine (50), L-proline (50), L-lysine (50), L-threonine (50), L-cystine (50), sodium phosphate (1000), sodium bicarbonate (750), ascorbic acid (300), magnesium aspartate (150), nicotinamide (30), d- alpha tocopheryl acetate (20), ferrous fumarate (20), alpha -lipoic acid (10), calcium pantothenate (5), riboflavin (3), thiamine (2), beta -carotene (750 mcg), biotin (5 mcg), cholecalciferol (5 mcg), cyanocobalamin (5 mcg) and flavor. He stated that he was not as severely affected by alcohol while drinking and did not experience a hangover the following morning.

MECHANISM OF ACTION - None given in the source material. USE - Used for prophylaxis and/or treatment of at least one symptoms caused or exacerbated by consumption of a toxic compound such as

ethanol.

ADVANTAGE - The composition enhances the metabolism of ethanol and inhibits some of the biochemical changes associated with ethanol and its by-products.

Dwg.0/0

CPI FILE SEGMENT: FIELD AVAILABILITY:

MANUAL CODES:

AB; DCN

CPI: B04-C02; B06-F03; B07-A02; B07-B03; B10-B02J; B10-C04D; B14-M01A

L137 ANSWER 23 OF 33

WPIDS COPYRIGHT 2005 THE THOMSON CORP on STN 2003-597592 [56] WPIDS

ACCESSION NUMBER: DOC. NO. CPI:

C2003-161827

TITLE:

Alleviating or reducing toxic, nutritional, and

metabolic distarbances associated with cancer and cancer

chemotherapy, comprises administering composition comprising riboflavin, effector of urea cycle,

and specified amino acids.

DERWENT CLASS:

INVENTOR (S):

PATENT ASSIGNEE(S):

COUNTRY COUNT: PATENT INFORMATION: B05

BURZYNSKI, S 🗷

(BURZ-I) BUKZYNSKI S R 103

Searched by Barb O Bryen, STIC

PAT	rent	NO		1	KINI	D DA	ATE		WI	EEK		LA	I	PG I	1IAN	II	PC						
	200																						
WO	200	3045	5372	2	A1	200	300	505	(20	0035	56)	El	Ŋ		A61	LKO.	31-1	195					
	RW:	ΑT	ΒE	BG	CH	CY	CZ	DΕ	DK	EΑ	$\mathbf{E}\mathbf{E}$	ES	FΙ	FR	GB	GH	GM	GR	IE	IT	KΕ	LS	LU
		MC	MW	ΜZ	NL	OA	PT	SD	SE	SK	\mathtt{SL}	SZ	TR	TZ	UG	ZM	zw						
	W:	ΑE	AG	AL	AM	ΑT	ΑU	ΑZ	BA	BB	BG	BR	BY	BZ	CA	CH	CN	CO	CR	CU	CZ	DE	DK
		DM	DZ	EC	EE	ES	FΙ	GB	GD	GE	GH	GM	HR	HU	ID	IL	IN	IS	JP	KE	KG	ΚP	KR
		ΚZ	LC	LK	LR	LS	LT	LU	LV	MA	MD	MG	MK	MN	MW	MX	MZ	NO	NZ	OM	PH	PL	PT
		RO	RU	SC	SD	SE	SG	SI	SK	\mathtt{SL}	ТJ	TM	TN	TR	TT	TZ	UA	UG	UZ	VC	VN	YU	za
		z_{M}	zw																				
ΑU	200	2352	2843	3	Α1	200	306	510	(20	004	19)				A61	LKO:	31-3	195					
ΕP	145	0783	1		A1	200	409	901	(20	004	57)	El	Ŋ		A61	LKO:	31-1	195					
	R:	AL	ΑT	BE	BG	CH	CY	CZ	DE	DK	EE	ES	FΙ	FR	GB	GR	ΙE	IT	LI	LT	LU	$rac{r}{\Lambda}$	MC
		MK	NL	PT	RO	SE	SI	SK	TR														
KR	200	406	556	5	Α	200	407	722	(20	047	74)				A61	LK03	31-5	525					
BR	200	2014	4430)	Α	200	41:	103	(20	0048	32)				A61	LKO:	31-3	195					

APPLICATION DETAILS:

PATENT NO	KIND	APPLICATION	DATE		
US 2003105104	A1	US 2001-995010	20011127		
WO 2003045372	A1	WO 2002-US37354	20021121		
AU 2002352843	A1	AU 2002-352843	20021121		
EP 1450781	A1	EP 2002-789801	20021121		
		WO 2002-US37354	20021121		
KR 2004065565	Α	KR 2004-707754	20040521		
BR 2002014430	Α	BR 2002-14430	20021121		
		WO 2002-US3735	20021121		

FILING DETAILS:

PATENT NO	KIND	PATENT NO
AU 2002352843	A1 Based on	WO 2003045372
EP 1450781	A1 Based on	WO 2003045372
BR 2002014430	A Based on	WO 2003045372

PRIORITY APPLN. INFO: US 2001-995010 20011127

INT. PATENT CLASSIF.:

MAIN:

A61K031-195; A61K031-525

SECONDARY:

A23L001-305; A23L001-3055; A61K031-185; A61K031-198; A61K031-1988; A61K031-5255; A61P041-00; A61P041-000

INDEX:

A61K031:198; A61K031:195; A61K031-525

BASIC ABSTRACT:

US2003105104 A UPAB: 20030903

NOVELTY - Alleviating or reducing the **toxic**, nutritional, and metabolic disturbances associated with cancer and cancer **chemotherapy** comprises administering to a patient a composition comprising **riboflavin**, effector of the urea cycle, and the amino acids **alanine**, **glycine**, **serine**, **taurine**, **threonine** and **valine**.

ACTIVITY - Cytostatic; Antiemetic; Endocrine-Gen.

MECHANISM OF ACTION - None given in the source material.

USE - For alleviating or reducing toxic, nutritional and metabolic disturbances, e.g. fatigue, and weakness associated with cancer and cancer chemotherapy. The method increases energy and has potential for decreasing the size of tumors within the patient.

Toxic effects which may be suffered by patients undergoing cancer chemotherapy include pancytopenia, alopecia,

nausea and vomiting.

Dwq.0/0

FILE SEGMENT: CPT

FIELD AVAILABILITY: AB; DCN

MANUAL CODES: CPI: B03-C; B07-D05; B10-A09B; B10-B02C; B14-E05;

B14-H01; B14-J05; B14-R02

L137 ANSWER 24 OF 33 WPIDS COPYRIGHT 2005 THE THOMSON CORP on STN

ACCESSION NUMBER: 2004-003557 [01] WPIDS

DOC. NO. CPI:

C2004-001624

TITLE:

Foodstuffs for preventing obesity and reducing triglyceride level and body fat, contain organic compound, which forms complex with zinc and source of

DERWENT CLASS:

B05 D13

PATENT ASSIGNEE(S): (ARIT-I) ARITA J

COUNTRY COUNT:

PATENT INFORMATION:

PATENT NO KIND DATE WEEK LA PG MAIN IPC JP 2003319760 A 20031111 (200401) * 7 A23L001-304

APPLICATION DETAILS:

PATENT NO KIND APPLICATION DATE JP 2003319760 A JP 2003-7830 20030116

PRIORITY APPLN. INFO: JP 2002-100643 20020226

INT. PATENT CLASSIF.:

MAIN: A23L001-304

A23L001-30; A23L001-305; A61K031-315; A61K033-30; SECONDARY: A61K047-12; A61K047-18; A61K047-20; A61K047-22;

A61K047-42; A61P003-04; A61P003-06; A61P003-10

BASIC ABSTRACT:

JP2003319760 A UPAB: 20040102

NOVELTY - Foodstuffs contain organic compound, which forms a complex with zinc and source of zinc.

ACTIVITY - Anorectic; Antilipemic; Antidiabetic; Cytostatic; Antiarteriosclerotic; Cardiovascular-Gen.; Antianginal; Hypotensive; Cardiant.

The bait agent was administered to KK-Ay mouse having symptoms of type 2 diabetes. The increase and decrease in body weight was evaluated and the results are shown in figure 1 and 2.

MECHANISM OF ACTION - None given.

USE - As health food for preventing obesity and reducing triglyceride level and body fat (claimed) and treating glucose tolerance disorder, diabetes, insulin resistance syndrome, polycystic ovary syndrome, hyper lipidemia, arteriosclerosis, cardiovascular disorder, hyperglycemia, angina, hypertension, cardiac failure and taste disorder.

ADVANTAGE - The foodstuff is stable and safe without producing any side effects.

DESCRIPTION OF DRAWING(S) - The figure shows a graph representing change in body weight after administered zinc sulfate and vitamin U. (Drawing includes non-English language text).

Dwg.1/14

FILE SEGMENT: CPI

FIELD AVAILABILITY: AB; GI; DCN

MANUAL CODES:

CPI: B05-A03A; B06-D01; B06-D09; B06-D17; B07-A02A; B07-D03; B07-D04; B07-D09; B10-A07; B10-A22;

09/955010 Page 74 Jones

B10-B01B; B10-B02D; B10-B02E; B10-B02H; B10-B02J; B10-C02; B10-C03; B10-C04E; B14-E12; B14-F01; B14-F01D; B14-F02; B14-F02B; B14-F06; B14-F07; B14-F09; B14-H01; B14-S04; D03-H01T2

L137 ANSWER 25 OF 33 WPIDS COPYRIGHT 2005 THE THOMSON CORP on STN

ACCESSION NUMBER: 2003-847970 [79] WPIDS DOC. NO. CPI: C2003-238964

Infusion solution for intravenous administration, TITLE:

comprises saccharide, amino acid and electrolyte.

DERWENT CLASS: B05 B07

PATENT ASSIGNEE(S): (AJIN) AJINOMOTO KK COUNTRY COUNT: 1

PATENT INFORMATION:

WEEK LA PG MAIN IPC PATENT NO KIND DATE _____ JP 2003252757 A 20030910 (200379)* 11 A61K031-198

APPLICATION DETAILS:

APPLICATION PATENT NO KIND _____ JP 2003252757 A JP 2002-377386 20021226

PRIORITY APPLN. INFO: JP 2001-399468 20011228

INT. PATENT CLASSIF.:

MAIN: A61K031-198

NDARY: A61K009-08; A61K031-07; A61K031-122; A61K031-197; SECONDARY:

A61K031-355; A61K031-375; A61K031-401; A61K031-405; A61K031-4172; A61K031-4188; A61K031-4415; A61K031-455; A61K031-51; A61K031-525; A61K031-59; A61K031-7004; A61K031-714; A61K033-14; A61K033-30; A61K033-42;

A61P003-02

BASIC ABSTRACT:

JP2003252757 A UPAB: 20031208

NOVELTY - Infusion solution for intravenous administration, comprises saccharide, amino acid and electrolyte. The infusion solution has total amount of nitrogen and nonprotein calorie amount ratio of 1:70-1:140 and has total energy of 450-625 kcal/l.

DETAILED DESCRIPTION - An INDEPENDENT CLAIM is also included for infusion solution kit, which contains infusion solution for intravenous administration and exclusive catheter.

ACTIVITY - Anabolic.

MECHANISM OF ACTION - None given.

USE - For intravenous administration to improve nutritional balance. ADVANTAGE - The infusion solution is safe with respect to patient at the time of moderate infestation and does not provide side

effects. The infusion solution improves the nutritional balance.

Dwq.0/0

CPI FILE SEGMENT: FIELD AVAILABILITY: AB; DCN

MANUAL CODES: CPI: B03-L; B05-A01A; B05-A01B; B05-A03A; B05-B02A3;

B05-C07; B06-D01; B06-D09; B07-D03; B07-D04C; B07-D09; B10-A04; B10-A07; B10-A17; B10-B01B;

B10-B02C; B10-C04D; B12-M07; B14-E11

L137 ANSWER 26 OF 33 WPIDS COPYRIGHT 2005 THE THOMSON CORP on STN

ACCESSION NUMBER: 2002-141648 [19] WPIDS

DOC. NO. CPI: C2002-043846

TITLE: Zinc-oligopeptide useful in the treatment of diabetes mellitus is prepared by proteolyzing a protein in water to obtain oligopeptides, which are further chelated with zinc ion.

DERWENT CLASS:

B02 D13

INVENTOR(S):

JI, S K; JIH, S G

PATENT ASSIGNEE(S):

(JISK-I) JI S K; (JIHS-I) JIH S G

COUNTRY COUNT:

30

PATENT INFORMATION:

PATENT NO K	KIND DATE	WEEK LA PG	MAIN IPC	X
		(200219)* EN 8		
R: AL AT BE RO SE SI		ES FI FR GB GR II	E IT LI LT LU L	V MC MK NL PT
US 2002028769	A1 20020307	(200221)	A61K038-16	
JP 2002034592	A 20020205	(200225)	5 C12P021-06	/
CN 1333372	A 20020130	(200231)	C12P021-02	
KR 2002006114	A 20020119	(200251)	C07K001-12	/ .
US 6740502	B2 20040525	(200435)	C12P021-06	
KR 421466	B 20040310	(200444)	C07K001-12	
US 2004126410	A1 20040701	(200444)	A61K038-16	

APPLICATION DETAILS:

PATENT NO	KIND	APPLICATION	DATE
EP 1172373	A2	EP 2001-710010	20010306
US 2002028769	A1	US 2000-730542	20001207
JP 2002034592	A	JP 2000-379180	20001213
CN 1333372	A	CN 2001-103060	20010122
KR 2002006114	A	KR 2000-39595	20000711
US 6740502	B2	US 2000-730542	20001207
KR 421466	В	KR 2000-39595	20000711
US 2004126410	A1 Div ex	US 2000-730542	20001207
		US 2003-734172	20031215

FILING DETAILS:

PATENT NO	KI	ND		PATENT	NO
		- 			
KP 421466	B	Previous	Publ Ki	200200	06114

PRIORITY APPLN. INFO: KR 2000-39595

20000711

INT. PATENT CLASSIF.:

MAIN: A61K038-16; C07K001-12; C07K007-06; C12P021-02;

C12P021-06

SECONDARY: A23J003-34; A23L001-30; A23L001-302; A23L001-304;

A23L001-305; A23L002-38; A23L002-52; A61K031-505;

A61K031-70; C07K014-415; C07K014-435

ADDITIONAL: A61K009-20; A61K009-48; A61K033-30; A61K047-42;

A61P003-02

BASIC ABSTRACT:

EP 1172373 A UPAB: 20020820

NOVELTY - In the preparation of a zinc-oligopeptide a suspension of protein is proteolyzed in water to form a mixture of oligopeptide, followed by chelating zinc ion with the oligopeptide.

DETAILED DESCRIPTION - Preparation of a zinc-oligopeptide involves: proteolyzing a suspension of protein in water at a pH of 6.8 - 9 in the presence of a protease to give a mixture of oligopeptides; and chelating zinc ions with the oligopeptides to obtain the zinc-oligopeptide solution.

INDEPENDENT CLAIMS are also included for the following:

1) zinc-oligopeptide of formula (I);

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Glu = glutamic acid;
         Asp = aspartic acid;
    Lys
         = lysine;
         Arg = arginine;
    Gly
         = glycine;
    Ala = alanine.
          2) a beverage comprising (I) in combination with at least one of a
    vitamin-C, vitamin-B1, vitamin-B2, fructose, alpha
     -amylase decomposed starch or magnesium stearate; and 3) a capsule or
     tablet prepared by dehydrating the beverage.
         ACTIVITY - Antidiabetic; cytostatic; vulnerary; antiseborrheic;
     dermatological; antirheumatic; antiarthritic; immunostimulant.
         MECHANISM OF ACTION - Insulin activator; mutant gene expression
     inhibitor; DNA polymerization promoter.
         USE - In the preparation of a beverage, which is further used in the
     preparation of a capsule or a tablet (claimed). (I) is useful in the
     treatment of diabetes mellitus, in anticancer therapy, and in
     the regeneration of injured tissue e.g. to accelerate wound healing,
    prevention of prostate problems, hair loss and treatment of acne and
     rheumatoid arthritis.
         ADVANTAGE - (I) has a molecular weight of 800 - 1,200 which, is
     smaller than the average molecular weight (24,000 - 28,000) of the
    membrane integral proteins of the small intestine, it can be readily
     absorbed by the body. (I) is water-soluble, thus its absorption by the
    body is not inhibited by the other compounds present in the digestive
     tract.
     Dwg.0/2
                      CPI
FILE SEGMENT:
FIELD AVAILABILITY:
                      AB; GI; DCN
                      CPI: B03-B; B03-C; B03-F; B04-C01B; B04-C02B2; B04-J03A;
MANUAL CODES:
                           B04-L05B; B04-N01A; B04-N02A; B05-A01B; B05-A03A;
                           B10-A07; B12-M11; B14-C06; B14-C09; B14-G01;
                           B14-H01B; B14-L06; B14-N17; B14-N17B; B14-N17D;
                           B14-S04; D03-H01G; D03-H01T2
L137 ANSWER 27 OF 33 WPIDS COPYRIGHT 2005 THE THOMSON CORP on STN
                     2002-025903 [03] WPIDS
ACCESSION NUMBER:
                     N2002-020018
DOC. NO. NON-CPI:
DOC. NO. CPI:
                     C2002-007233
                      Detecting specific enzyme activities, useful e.g. for
TITLE:
                      identifying enzyme inhibitors, based on enzymatic
                      conversion of precursor to factor for which host cell is
                      auxotrophic.
                      B04 C06 D16 S03
DERWENT CLASS:
INVENTOR(S):
                      SILVA, C J
                      (CUBI-N) CUBIST PHARM INC
PATENT ASSIGNEE(S):
COUNTRY COUNT:
                      95
PATENT INFORMATION:
                  KIND DATE
                                  WEEK
                                               PG MAIN IPC
     PATENT NO
                                          LA
     WO 2001077366 A1 20011018 (200203)* EN 51 C12Q001-00
       RW: AT BE CH CY DE DK EA ES FI FR GB GH GM GR IE IT KE LS LU MC MW MZ
            NL OA PT SD SE SL SZ TR TZ UG ZW
        W: AE AG AL AM AT AU AZ BA BB BG BR BY BZ CA CH CN CO CR CU CZ DE DK
            DM DZ EE ES FI GB GD GE GH GM HR HU ID IL IN IS JP KE KG KP KR KZ
            LC LK LR LS LT LU LV MA MD MG MK MN MW MX MZ NO NZ PL PT RO RU SD
            SE SG SI SK SL TJ TM TR TT TZ UA UG US UZ VN YU ZA ZW
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AU 2001057002

C12Q001-00

A 20011023 (200213)

PATENT NO	KIND	APPLICATION	DATE
WO 2001077366	A1	WO 2001-US11567	20010410
AU 2001057002	Α	AU 2001-57002	20010410

FILING DETAILS:

PRIORITY APPLN. INFO: US 2000-195911P 20000410

INT. PATENT CLASSIF.:

MAIN: C12Q001-00

SECONDARY: C12N015-00; C12N015-63; G01N033-573

BASIC ABSTRACT:

WO 200177366 A UPAB: 20020114

NOVELTY - Detecting a particular enzymatic activity (A) by treating host cells (B), genetically engineered to express at least one activity, with a precursor (C) of a factor (C') for which the cells are auxotrophic. (C) is converted only if the cells express (A), so if (B) survive when cultured under auxotrophic conditions this indicates expression of (A).

DETAILED DESCRIPTION - INDEPENDENT CLAIMS are also included for the following:

- (1) similar method in which cells are treated with (C) before transformation to express (A);
- (2) method for producing a protein (I) by introducing into a host cell a gene encoding (I) and a gene encoding an enzyme, then culturing cells under auxotrophic conditions in presence of (C) so that only cells that express both genes can grow;
- (3) replicable vector containing a gene that expresses (A) able to convert (C) to (C'), thus allowing growth of transformed cells under auxotrophic conditions in presence of (C);
 - (4) host cell, auxotrophic for (C), that expresses (A);
- (5) kit comprising, in separate vessels, auxotrophic cells and a replicable vector; and
 - (6) a method for detecting an enzyme inhibitor.

USE - The method is used for identifying new enzymes with specific activities or enzymatic pathways, e.g. in a gene expression library, selecting host cells, maintaining plasmids without use of antibiotics, expressing proteins and identifying specific enzyme inhibitors.

ADVANTAGE - The method produces only positive results, does not require chromophores or fluorophores, uses substrates that are more like their targets, and is accurate and efficient with 1 million or more assays being done in a single petri dish overnight. Since selection is based on auxotrophy and synthetic non-toxic compounds (not on antibiotics), (B) may be acceptable for release into the environment. (C) are generally more stable than antibiotics, so should be suitable for organisms that grow in extreme environments.

Dwg.0/2

FILE SEGMENT: CPI EPI FIELD AVAILABILITY: AB; DCN

MANUAL CODES: CPI: B01-D02; B03-A; B03-B; B03-C; B03-G; B04-B01B;

B04-B03A; B04-B03B; B04-C03; B04-D01; B04-E01; B04-E08; B04-F0100E; B04-L01; B04-L02; B04-N0400E; B05-B01P; B06-D01; B06-D02; B06-D05; B06-D09; B06-D18; B06-F03; B07-A01; B07-B03; B07-D03; B07-D04C; B07-D09; B10-A17; B10-A22; B10-B02; B10-B03B; B10-C02; B10-C04A; B10-C04E; B10-D02; B10-E04D; B10-J02; B11-C08E3; B12-K04; C01-D02; C03-A; C03-B; C03-C; C03-G; C04-B01B; C04-B03A;

Page 78 09/955010 Jones

C04-B03B; C04-C03; C04-D01; C04-E01; C04-E08; C04-F0100E; C04-L01; C04-L02; C04-N0400E; C05-B01P; C06-D01; C06-D02; C06-D05; C06-D09; C06-D18; C06-F03; C07-A01; C07-B03; C07-D03; C07-D04C; C07-D09; C10-A17; C10-A22; C10-B02; C10-B03B; C10-C02; C10-C04A; C10-C04E; C10-D02; C10-J02; C11-C08E3; C12-K04; D05-A02A; D05-A02B; D05-A02C; D05-A02D; D05-A02E; D05-A02F; D05-H09; D05-H12E; D05-H14

EPI: S03-E14H4

L137 ANSWER 28 OF 33 WPIDS COPYRIGHT 2005 THE THOMSON CORP on STN

ACCESSION NUMBER:

2001-602729 [68] WPIDS

DOC. NO. CPI:

C2001-178562

TITLE:

Production of standardized drinks and potable water for health profiles of nutritional needs, involves dissolving

specific additives into the drinks or water.

DERWENT CLASS:

B05 D13 D15 D16

INVENTOR (S):

COSTA, F

PATENT ASSIGNEE(S):

(DCOS-I) MOREIRA DA COSTA F J; (COST-I) COSTA F

COUNTRY COUNT:

PATENT INFORMATION:

PATENT NO					
WO 2001068534					
RW: AT BE CH					-
	CN IL IN IS				
PT 102430					
AU 2001041301					
EP 1307408	A1 20030507	(200332)	EN C	02F001-68	
				I LU MC NL SE	TR
US 2004013784	A1 20040122	(200407)	C:	12C001-00	

APPLICATION DETAILS:

PATENT NO	KIND	APPLICATION	DATE
WO 2001068534	A1	WO 2001-PT3	20010315
PT 102430	A	PT 2000-102430	20000316
AU 2001041301	A	AU 2001-41301	20010315
EP 1307408	A1	EP 2001-912612	20010315
		WO 2001-PT3	20010315
US 2004013784	A1	WO 2001-PT3	20010315
		US 2003-239621	20030127

FILING DETAILS:

PATENT NO	KIND	PATENT NO
AU 2001041301	A Based on	WO 2001068534
EP 1307408	Al Based on	WO 2001068534

PRIORITY APPLN. INFO: PT 2000-102430 20000316

INT. PATENT CLASSIF.:

MAIN: C02F001-68; C12C001-00

SECONDARY: A23L001-29

BASIC ABSTRACT:

WO 200168534 A UPAB: 20011121

NOVELTY - Production of standardized drinks and potable water comprises dissolving specifically produced additives, in the form of solids (tablets or gelatinous capsules), liquids, gases, and/or energy, into the drinks or

water (distilled or demineralized). The additives are elements and chemical compounds needed for daily human consumption, e.g. vitamins, amino acid, proteins, minerals.

USE - The invention produces standardized drinks and potable water for health profiles of nutritional needs. The drinks can be milk, drinkable yogurt, coffee, tea, chocolate, shake, juice, soda, sparkling canned drink, beer, wine, liquor, brandy, whiskey, spirituous drink, white drink, and other alcoholic and non-alcoholic drink. The water is from natural resources, e.g. water springs, fountains, rivers, or from artificial resources, e.g. public water networks, holes, and bottled waters.

ADVANTAGE - The invention provides high quality drinks and potable water regionally standardized for specific and/or pre-defined health profiles of the consumers. The use of the additives in the production of standardized drinks and potable water satisfies the biological nutritional needs. Some illnesses, caused by many factors including lack of nutrients, and conditions that the invention will satisfy are uric acid - gout; alcoholism; allergies; anemia; anorexia; arterioscleroses; arthritis; renal calculus in kidneys and vesicle; cancer; growth; high cholesterol; depression; dehydration; malnutrition; hormone clutter; sports - athletes; diabetes mellitus; genetic disorders; psychological illnesses; psychiatric illnesses; bone diseases; ear pain; skin diseases; alimentary and digestive system illnesses; cardiovascular system illnesses; sanguineous circulatory system illnesses; hepatic system illnesses; immunology system illnesses; lymphatic system illnesses; nervous system illnesses; ophthalmologic system illnesses; respiratory system illnesses; reproductive system illnesses; urinary and renal system illnesses; fatique; infertility; human immunodeficiency virus (HIV)/acquired immune deficiency syndrome (AIDS); leukemia; maternity; menopause; obesity; corporal odor; osteoporosis; high arterial pressure; arterial pressure decrease; menstrual problems; prostate; puberty; kidneys; rheumatism; toxic-dependence - drugs; ulcer; oldness; vesicle. Dwg.0/0

FILE SEGMENT: CPI
FIELD AVAILABILITY: AB; DCN

MANUAL CODES: CPI: B03-L; B04-A10; B04-L01; B04-N04; B05-A01A;

B05-A01B; B05-A03; B05-B02A3; B05-B02C; B05-C07; B06-H; B07-H; B10-A06; B10-A22; B10-B02A; B10-B02B;

B10-C04D; B14-E11; D03-B08; D03-D01; D03-D02; D03-H01F; D03-H01G; D03-H01H; D03-H01T2; D04-A;

D05-E

L137 ANSWER 29 OF 33 WPIDS COPYRIGHT 2005 THE THOMSON CORP on STN

ACCESSION NUMBER: 2001-328414 [34] WPIDS

DOC. NO. CPI: C2001-100693

TITLE: Treating neurobehavioral disorders comprises

administering a composition comprising amino acid(s) and e.g. vitamins, neurotransmitter precursors, minerals, corticosteroids, enzyme inhibitors and/or immunological

enhancers.

DERWENT CLASS: B05

INVENTOR(S): BECHTHOLD, J C; LILLY, T D

PATENT ASSIGNEE(S): (BECH-I) BECHTHOLD J C; (LILL-I) LILLY T D

COUNTRY COUNT: 9

PATENT INFORMATION:

PATENT NO KIND DATE WEEK LA PG MAIN IPC

WO 2001026642 A2 20010419 (200134) * EN 91 A61K031-00

RW: AT BE CH CY DE DK EA ES FI FR GB GH GM GR IE IT KE LS LU MC MW MZ

NL OA PT SD SE SL SZ TZ UG ZW

W: AE AG AL AM AT AU AZ BA BB BG BR BY BZ CA CH CN CR CU CZ DE DK DZ

EE ES FI GB GD GE GH GM HU ID IL IN IS JP KE KG KP KR KZ LC LK LR LS LT LU LV MA MD MG MK MN MW MX MZ NO NZ PL PT RO RU SD SE SG SI SK SL TJ TM TR TT TZ UA UG UZ VN YU ZA ZW

AU 2000080038 A 20010423 (200147)

A61K031-00

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APPLICATION DETAILS:

PATENT NO	KIND	APPLICATION	DATE
WO 2001026642	A2	WO 2000-US27894	20001006
AU 2000080038	A	AU 2000-80038	20001006

FILING DETAILS:

PATENT NO	KIND	PATENT NO
AU 2000080038	A Based on	WO 2001026642

PRIORITY APPLN. INFO: US 2000-201043P

US 2000-201043P 20000501; US 1999-158604P 19991008; US 1999-166068P 19991117

INT. PATENT CLASSIF.:

MAIN:

A61K031-00

BASIC ABSTRACT:

WO 200126642 A UPAB: 20010620

NOVELTY = Treating a neurobehavioral disorder comprises administering intravenously a composition comprising amino acid(s), neurotransmitter precursors, vitamins, inhibitors of neurotransmitter degradation and/or immune function enhancers.

DETAILED DESCRIPTION - Treating a neurobehavioral disorder comprises administering intravenously a composition comprising amino acid(s), neurotransmitter precursors, vitamins, inhibitors of neurotransmitter degradation and/or immune function enhancers.

INDEPENDENT CLAIMS are included for:

- (1) a sterile composition (I) for treating neurobehavioral disorders comprising:
 - (a) at least one amino acid;
 - (b) vitamin C; and
 - (c) an electrolyte solution.
- (2) a sterile composition (II) for treating neurobehavioral disorders comprising:
 - (a) at least one amino acid;
 - (b) a corticosteroid; and
 - (c) an electrolyte solution;
- (3) a sterile composition (III) for treating neurobehavioral disorders comprising:
- (a) vitamin C;
 - (b) a corticosteroid; and
 - (c) an electrolyte solution;
- (4) a sterile composition (IV) for treating neurobehavioral disorders comprising:
 - (a) at least one amino acid;
 - (b) an immune potentiating amount of gamma-globulin; and
 - (c) an electrolyte solution;
- (5) a sterile composition (V) for treating neurobehavioral disorders comprising:
 - (a) at least one amino acid;
 - (b) an inhibitor of opioid peptide degradation; and
 - (c) an electrolyte solution;
- (6) an oral composition (VI) for treating neurobehavioral disorders comprising:

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- (a) at least one amino acid; and
- (b) a substance selected from Ginko Biloba, methylsulfonylmethane, phosphatidylserine, phosphatidylcholine, alpha lipoic acid, red ginseng root, L-aspartic acid, ephedrine, pancreatic enzymes, caffeine, theobromine, Hypericum perforatum extract, S-adenosyl methionine, dihydroxyacetate, DMAE, grape seed extract, betaine, prickly pear cactus extract, Gymnea sylvestre extract, nicotinamide adenine dinucleotide/hydrogen, cholecystokinin, Cyclo (His-Pro), corticotropin-releasing hormone, neuropeptide Y, galanin, monolaurin or fructo-oligosaccharides.
- (7) a method for treating a neurobehavioral disorder comprising administering intravenously a sterile and isotonic composition comprising:(a) vitamin C;
 - (b) a corticosteroid; and
- (c) water;
 - (8) a method for treating a neurobehavioral disorder comprising:
 - (i) evaluating a neurobiological characteristic of the disorder; and
- (ii) injecting the patient with an intravenous composition to treat the disorder; and
- (9) a composition (VII) for treating a neurobehavioral disorder comprising:
 - (i) an inhibitor of opioid degradation; and
- (ii) a substance selected from group (A) which comprises thymus extract, L-taurine, alpha-keto glutarate, lidocaine, L-glutathione, pyridoxal-5-phosphate, sodium ascorbate, oxytocin, L-glycine, L-leucine, gamma globulin, vitamin B complex, magnesium taurate, citric acid, chromium polynicotinate, chromium nicotinate, chromium picolynate, zinc chelate, calcium chelate, vitamin B-12, vitamin B-5, vitamin B-6, vitamin B-1, folic acid, L-taurine and balanced amino acid solution with electrolytes.

ACTIVITY - Anti-alcoholic, anti-depressant; nootropic; antismoking; antiaddictive; anxiolytic; tranquilizer; anorectic; neuroleptic; anticonvulsant; neuroprotective.

A 38 year old male suffering from sleep disorders, obsessive-compulsive disorder, anger and rage disorder, depression, drug and alcohol addiction, attention deficit hyperactivity disorder, neurally mediated hypotension, chronic fatigue syndrome, dyslexia and a history of debilitating brain disorder for whom conventional therapies had minimal effect was given a number of infusion treatments culminating in an infusion comprising saline (500 ml), sodium ascorbate (25 mg), molybdenum (250 mg), magnesium (600 mg), vitamin E (500 IU), vitamin B1 + B complex (1 cc), manganese (2 cc), zinc (1 cc), selenium (2 cc), chromium (2 cc), calcium gluconate (7 cc), taurine (2 cc), copper solution (2 cc), adrenal cortical extract (5 cc) and vitamin A (1000000 IU). The subject noted a reduction in craving, fluid retention was improved and blood pressure stabilized. The subject also experienced an increased sense of calm and increased motivation, mood and energy.

MECHANISM OF ACTION - The components of the compositions are e.g. enzyme inhibitors (for inhibiting neurotransmitter degradation or opiate degradation), neurotransmitter precursors, insulin potentiators, dopamine receptor agonists, opiate receptor antagonists and ammonia scavengers.

USE - The compositions are useful for reducing symptoms associated with withdrawal, improving symptoms of drug and alcohol overuse and reducing or preventing cravings for addictive substances. The compositions and methods permit the brain to function more normally by supporting or increasing the function of deficient neurochemical pathways and can eliminate or decrease symptoms of withdrawal, craving or compulsion associated with addiction and other central neurobiological disorders. The methods are useful for treating neurobehavioral disorders and for diagnosing and/or evaluating underlying neurobehavioral disorders. The treatments are also useful for disorders involving carbohydrate addiction, weight gain and nicotine addiction. Neurobehaviors treatable by these

methods and compositions include e.g. obesity, smoking, Tourette's Syndrome, ADHD (attention deficit hyperactivity disorders), ADD (attention deficit disorders), Schizoid/Avoidant Behavior, aggression, posttraumatic stress syndrome, alcoholism, drug addiction, obsessive compulsive disorders, learning disorders, reading problems, gambling, manic symptoms, phobias, panic attacks, oppositional defiant behavior, conduct disorder, sexual behavior disorders, schizoid disorders, somatization disorders, depression, sleep disorders, general anxiety disorders, stuttering, tic disorders, anger and violent behavior disorders as well as Huntington's chorea, amyotropic lateral sclerosis, environmental sensitivity, chemical injury syndrome and chronic fatigue syndrome.

ADVANTAGE - The compositions can minimize adverse effects of addiction and other neurobehavioral disorders in patients recovering from these disorders. The compositions can eliminate or decrease symptoms of withdrawal, craving or compulsion associated with addiction and other central neurobiological disorders and these effects can result in longer lasting improvements in symptoms, thus reducing the risk of relapse and also making it more likely that the patient will complete their course of treatment. The compositions are less expensive in comparison to the current costs of residential treatment for drug and alcohol addiction and costs incurred due to repeat therapy can be reduced.

Dwg.0/0

FILE SEGMENT: CPI

FIELD AVAILABILITY: AB; DCN

MANUAL CODES:

CPI: B03-B; B03-D; B03-F; B03-H; B03-L; B04-B04G; B04-C01B; B05-A01B; B05-A03A; B05-A03B; B06-D01; B06-D09; B07-D04C; B10-B02; B10-B02B; B10-B02E; B10-C04E; B10-G02; B14-E12; B14-J01; B14-J01A1; B14-J01A4; B14-J01B2; B14-J01B3; B14-J01B4; B14-J07; B14-M01; B14-M01B

L137 ANSWER 30 OF 33 WPIDS COPYRIGHT 2005 THE THOMSON CORP on STN

ACCESSION NUMBER: 2002-084181 [12] WPIDS

DOC. NO. CPI: C2002-025785

TITLE:

Micronutrient combination product based on vitamins,

folic acid, magnesium, arginine, coenzyme Q10,

carotenoids and omega fatty acids, useful as nutritional

supplement in drug treatment and smoking.

DERWENT CLASS:

PATENT ASSIGNEE(S):

(ORTH-N) ORTHOMOL PHARM VERTRIEBS GMBH

COUNTRY COUNT:

PATENT INFORMATION:

PATENT NO KIND DATE WEEK LA PG MAIN IPC _____ DE 20116346 U1 20011220 (200212)* 20 A61K031-505

APPLICATION DETAILS:

APPLICATION DATE PATENT NO KIND _____ DE 2001-20116346 20011005 DE 20116346 U1

PRIORITY APPLN. INFO: DE 2001-20116346 20011005

INT. PATENT CLASSIF.:

MAIN: A61K031-505 SECONDARY: A61K031-355

BASIC ABSTRACT:

DE 20116346 U UPAB: 20020221

NOVELTY - A micronutrient combination product comprises vitamin C, vitamin E, vitamin B6, vitamin B12, folic acid, magnesium, arginine,

09/955010 Page 83 Jones

coenzyme Q10, carotenoid and omega fatty acids, each in a daily dosage range.

DETAILED DESCRIPTION - The micronutrient combination preparation comprises, on a daily dose basis:

- (a) vitamin C (350-750 mg, preferably 450-650 mg, especially 500-600 mg, particularly 520-560 mg);
- (b) vitamin E (100-200 mg, preferably 120-180 mg, especially 130-270 mg, particularly 140-160 mg);
- (c) vitamin B6 (3-25 mg, preferably 4-20 mg, especially 5-18 mg, particularly 5-15 mg);
 - (d) vitamin B12 (6-12 mg, preferably 7-11 mg, especially 8-9 mg);
- (e) folic acid (400-1,000 micro g, preferably 450-900 micro g, especially 500-850 micro g, particularly 600-800 micro g);
- (f) magnesium (150-300 mg, preferably 160-250 mg, especially 170-220 mg, particularly 180-200 mg);
- (g) arginine (100-400 mg, preferably 110-300 mg, especially 120-250 mg, particularly 125-200 mg);
- (h) coenzyme Q10 (10-20 mg, preferably 11-19 mg, especially 12-18 mg, particularly 15-16 mg);
- (i) carotenoid (2-10 mg, preferably 3-9 mg, especially 4-8 mg, particularly 5-6 mg); and
- (j) omega-3-fatty acids (400-1000 mg, preferably 420-900 mg, especially 450-800 mg, particularly 500-600 mg).

ACTIVITY - None given in the source material.

MECHANISM OF ACTION - None given in the source material.

USE - The product is used to prevent nutritional deficiency disorders, caused especially by drug treatment of human diseases, especially cardiac diseases, or by smoking.

ADVANTAGE - The product has no harmful side effects and can be safely administered together with drugs where it reduces their side effects.

Dwg.0/0

CPI FILE SEGMENT: FIELD AVAILABILITY: AB; DCN

MANUAL CODES:

CPI: B03-A; B03-B; B03-C; B03-D; B03-E; B03-F; B03-H; B03-K; B04-L02; B05-A01B; B05-A03; B05-B02C; B06-D09; B07-D04; B10-B02D; B10-B02J; B10-C04D; B10-C04E; B14-E11

L137 ANSWER 31 OF 33 WPIDS COPYRIGHT 2005 THE THOMSON CORP on STN

ACCESSION NUMBER:

1998-315266 [28] WPIDS

DOC. NO. CPI:

C1998-097296

TITLE:

Integumentary cortical function enhancer and infection

preventing composition - contains extract of wild

Tiliaceae annual herb, natural enzyme liquor, pure water, saccharide, licensed food additive, cosmetic raw material

and pharmaceutical.

DERWENT CLASS:

A96 B04

PATENT ASSIGNEE(S):

(SAKA-I) SAKATA S

COUNTRY COUNT:

PATENT INFORMATION:

LA PG MAIN IPC PATENT NO KIND DATE WEEK JP 09176029 A 19970708 (199828)* 3 A61K035-78

APPLICATION DETAILS:

PATENT NO	KIND	APPLICATION	DATE
JP 09176029	Α	JP 1995-354770	19951226

PRIORITY APPLN. INFO: JP 1995-354770 19951226

INT. PATENT CLASSIF.:

MAIN: A61K035-78

SECONDARY: A61K031-70; A61K031-715; A61K038-43

BASIC ABSTRACT:

JP 09176029 A UPAB: 19980715

Integumentary cortical function enhancer and infection preventing composition contains extract of wild Tiliaceae annual herb, naturally occurring enzyme liquor, pure water, saccharide, licensed food additive, standard cosmetic raw material and licensed pharmaceutical.

The extract of wild Tiliaceae annual herb preferably contains at least 1 of protein, lipid, ash, carbohydrate, phosphorus, iron, calcium, potassium, carotene, vitamin-A, thiamine, riboflavin, ascorbic acid and water. The naturally occurring enzyme liquor includes at at least 1 of amino acid e.g. arginine, lysine, histidine, phenylalanine, tyrosine, leucine, isoleucine, methionine, valine,

alanine, glycine, proline, glutamine, serine,

threonine, aspartic acid, tryptophane, cystine, vitamin-A, -B and
-E, and pure water.

ADVANTAGE - The composition is a safe alternative agent to a steroidal composition and has no side effects.

Dwg.0/0

FILE SEGMENT: CPI FIELD AVAILABILITY: AB; DCN

MANUAL CODES: CPI: A12-V01; B04-A08C2; B04-A10; B04-C02; B04-D01;

B04-L01; B14-D01D; B14-E11; B14-R01

L137 ANSWER 32 OF 33 WPIDS COPYRIGHT 2005 THE THOMSON CORP on STN

ACCESSION NUMBER:

1992-433373 [52] WPIDS

DOC. NO. CPI:

C1992-192375

TITLE:

Nutrient compsn. for treating immune disorders e.g. AIDS - including (non)oxidised gamma-L-glutamyl-L-cysteinyl

glycine, gamma-L-glutamyl-L-cysteine etc. used

with antiviral drugs.

DERWENT CLASS:

B04 B05

INVENTOR(S):

MAHNAZ KHALED, F; MAHNAZ, K F; KHALED, F M

PATENT ASSIGNEE(S):

(LIFE-N) LIFE SCI TECHNOLOGIES INC

COUNTRY COUNT:

PATENT INFORMATION:

PAT	TENT NO	KIN	D DATE	WEEK	LA PG	MAIN IPC	
WO						A61K037-02	
	RW: AT BE	CH DE	DK ES FR	GB GR IT	LU MC NL	SE	
	W: AU BB	BG BR	CA FI HU	JP KP KR	LK MG MW	NO PL RO RU S	SD US
ΑU	9221879	Α	19930108	(199315)		A61K037-02	
EP	604433	A1	19940706	(199426)	EN	A61K037-02	
	R: DE FR	GB					
EP	604433	A4	19941012	(199534)		A61K037-02	
US	5977073	Α	19991102	(199953)		A61K038-00	
ΕP	604433	B1	20000315	(200018)	EN	A61K038-00	
	R: DE FR	GB					
DE	69230796	E	20000420	(200026)		A61K038-00	

APPLICATION DETAILS:

PATENT NO	KIND	APPLICATION	DATE
WO 9221368	A1	WO 1992-US4653	19920604
AU 9221879	* A	AU 1992-21879	19920604 19920604
		WO 1992-US4653	1992060

E	EP 604	433	A1	ΕP	1992-913917	19920604
				WO	1992-US4653	19920604
E	EP 604	433	A4	EP	1992-913917	
τ	JS 5 97	7073	A	US	1991-711530	19910606
E	EP 604	433	B1	ΕP	1992-913917	19920604
				WO	1992-US4653	19920604
Ι	E 692	30796	E	DE	1992-630796	19920604
				EΡ	1992-913917	19920604
				WO	1992-US4653	19920604

FILING DETAILS:

PATENT NO	KIND	PATENT NO
AU 9221879	A Based on	WO 9221368
EP 604433	Al Based on	WO 9221368
EP 604433	B1 Based on	WO 9221368
DE 69230796	E Based on	EP 604433
	Based on	WO 9221368

PRIORITY APPLN. INFO: US 1991-711530 19910606

REFERENCE PATENTS: 2.Jnl.Ref; US 4466978; US 4927808; WO 9102535

INT. PATENT CLASSIF.:

MAIN: A61K037-02; A61K038-00

SECONDARY: C07K005-06; C07K005-08

BASIC ABSTRACT:

WO 9221368 A UPAB: 19931118

Compsn. for treating immune disorders in mammals comprises (1) 50-3000 mg. of a cpd. (I) or its salt or ester, which directly enhances the level of gamma-L-glutamyl-L- cysteinylglycine (Ia), (2) 50-3000 mg. L-glutamine, (3) 50-1000 mg. Vitamin C, (4) 50-500 mg. Vitamin E, (5) 10-100 mg. beta-carotene, and (6) 1-25 mg. Vitamin B6, all components being in purified form.

(I) is specifically (Ia), gamma-L-glutamyl-L-cysteine,

N-acetyl-L-cysteine or N-acetyl-L-cysteinyl-glycine.

The compsn. also contains at least one (pref. all) of 50-5000 mg. L-arginine, 5-50 micro-g. Cr, 50-150 micro-g. folic acid, 1-5 mg. Fe, 10-50 mg. Mg, 5-50 mg. pantothenic acid, 1-2.5 mg. riboflavin, 5-50 mg. thiamine, 0.5-10 mg. Vitamin A, 10-1000 micro-g. Se, 0.5-5 micro-g. Vitamin B12 and 1-50 mg. Zn.

USE/ADVANTAGE - The compsn. improves immune competence of the patient. It is especially used where immune deficiency is caused by a virus or bacterium and the patient is being treated with an appropriate anti-organism agent which has some **toxicity** for the host. In such cases the compsn. reduces **toxicity** and accelerates replication of the pathogen, making it more susceptible to the agent. Dwg.0/0

FILE SEGMENT: CPI FIELD AVAILABILITY: AB; DCN

MANUAL CODES: CPI: B03-L; B05-A01B; B05-A03; B05-B02C; B06-D09;

B10-A17; B10-B02D; B10-B02J; B10-C04D; B12-A01;

B12-A06; B12-D02A

L137 ANSWER 33 OF 33 WPIDS COPYRIGHT 2005 THE THOMSON CORP on STN

ACCESSION NUMBER: 1990-304797 [40] WPIDS

DOC. NO. CPI: C1990-131625

TITLE: Cellular growth medium - which allows maintenance and

propagation of cells in normal atmospheric carbon di

oxide.

DERWENT CLASS: B04 D16

INVENTOR(S): BOYD, M R; MONKS, A P; SCUDIERO, D A; SKEHAN, P J;

VISTICA, D T; MONKS, P A; VISTICA, A

PATENT ASSIGNEE(S):

(USDC) US DEPT OF COMMERCE; (USSH) NAT INST OF HEALTH;

(USDC) US SEC OF COMMERCE

COUNTRY COUNT:

PATENT INFORMATION:

PAT	ENT NO	KI	ND D	ATE		WE	EK		LA	I	P.G	MAIN	IPC
US	467939	A	.0 19	9009	04	(19	904	0)*	,				
WO	9110726	Z	. 19	9107	25	(19	913	2)					
	RW: AT BE	CH D	E DK	ES	FR	GB	GR	IT	LU	NL	SE		
	W: AU CA	JP											
ΑU	9173001	7	. 19	9108	05	(19	914	5)					
EP	512066	7	1 19	9211	11	(19	924	6)	EN	1	41	C121	1005-00
	R: AT BE	CH D	E DK	ES	FR	GB	GR	ΙT	LI	LU	NL	SE	
JP	05502379	W	19	9304	28	(19	932	2)			14	C121	1005-08
ΑU	653927	Е	19	9410	20	(19	944	3)				C121	1005-02
EP	512066	A	4 19	9304	28	(19	952	6)					
JP	07089954	Е	2 19	9510	04	(19	954	4)			23	C12(001-02
EP	512066	Е	1 19	9611	13	(19	965	0)	EN	1	38	C121	1005-00
	R: AT BE	CH D	E DK	ES	FR	GB	GR	IT	LI	LU	NL	SE	
DĒ	69123140	E	19	9612	19	(19	970	5)				C121	1005-00
CA	2074363	C	20	0411	09	(20	047	4)	EN	ı		C121	1005-02

APPLICATION DETAILS:

PATENT NO	KIND	APPLICATION	DATE
US 467939	A0	US 1990-213786	19900122
EP 512066	. A1	EP 1991-904443	19910122
		WO 1991-US451	19910122
JP 05502379	W	JP 1991-504839	19910122
		WO 1991-US451	19910122
AU 653927	В	AU 1991-73001	19910122
EP 512066	A4	EP 1991-904443	
JP 07089954	B2	JP 1991-504839	19910122
		WO 1991-US451	19910122
EP 512066	B1	EP 1991-904443	19910122
		WO 1991-US451	19910122
DE 69123140	E	DE 1991-623140	19910122
		EP 1991-904443	19910122
		WO 1991-US451	19910122
CA 2074363	С	CA 1991-2074363	19910122
		WO 1991-US451	19910122

FILING DETAILS:

PATENT NO	KIND	PATENT NO		
EP 512066 JP 05502379 AU 653927	A1 Based on W Based on B Previous Publ. Based on	WO 9110726 WO 9110726 AU 9173001 WO 9110726		
JP 07089954	Based on Based on	JP 05502379 WO 9110726		
EP 512066 DE 69123140	B1 Based on E Based on	WO 9110726 EP 512066		
CA 2074363	Based on C Based on	WO 9110726 WO 9110726		

PRIORITY APPLN. INFO: US 1990-213786 19900122; US 1990-467939 19900122

REFERENCE PATENTS: 2.Jnl.Ref; US 3883393; US 4411990; US 4533637; 3.Jnl.Ref;

9.Jnl.Ref

INT. PATENT CLASSIF.: C12N000-01; C12N005-02; C12Q001-02

MAIN: C12N005-00; C12N005-02; C12N005-08; C12Q001-02

SECONDARY: C12N000-01; C12Q001-04

ADDITIONAL: C12N005-06

INDEX: C12Q001-02, C12R001:91

BASIC ABSTRACT:

US N7467939 N UPAB: 20011211

Disclosed is a growth medium which allows maintenance and propagation of cells in normal atmospheric CO2, that is without requiring an exogenous, regulated supply of enriched CO2 for buffering.

The medium, termed PDRG basal growth medium, comprises 78 mg/l L-alanine, 265 mg/l L-arginine, 88 mg/l L-asparagine, 5 mg/l L-aspartic acid, 52 mg/l L-cysteine, 24 mg/l L-cysteine, 2HCl, 5 mg/l L-glutamic acid, 343 mg/l L-glutamine, 79 mg/l glycine, 141 mg/l L-histidine, 78 mg/l L-isoleucine, etc. USE/ADVANTAGE - The medium has good pH stability and buffering capacity in atmospheric CO2. It exhibits the capability for large scale growth of a broad range of human tumour cell lines for in vitro anticancer drug screening. @(33pp Dwg.No.0/7)

FILE SEGMENT: CPI FIELD AVAILABILITY: AB; DCN

MANUAL CODES: CPI: B03-B; B03-C; B03-E; B04-A06; B05-A01A; B05-A01B;

B05-A03A; B05-B02A3; B06-D09; B06-F03; B07-B03; B07-D04C; B07-D12; B10-A07; B10-A22; B10-B01B; B10-B02C; B10-C04E; B10-E04C; B10-E04D; D05-H01

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